

# Global Practices and Preferences in the Use of Osteobiologics for Anterior Cervical Discectomy and Fusion: A Cross-Sectional Study

Global Spine Journal  
2025, Vol. 0(0) 1–11  
© The Author(s) 2025  
Article reuse guidelines:  
[sagepub.com/journals-permissions](https://sagepub.com/journals-permissions)  
DOI: 10.1177/21925682251322417  
[journals.sagepub.com/home/gsj](https://journals.sagepub.com/home/gsj)



Luca Ambrosio, MD<sup>1,2</sup> , Arun Kumar Viswanadha, MS<sup>3</sup> ,  
Pieter-Paul A. Vergroesen, MD, PhD, MBA<sup>4</sup>, Zorica Buser, MBA, PhD<sup>5</sup> ,  
Hans Joerg Meisel, MD, PhD<sup>6</sup> , Nancy Santesso, PhD<sup>7</sup>, Jason P. Y. Cheung, MD<sup>8</sup>,  
Hai V. Le, MD<sup>9</sup> , Gianluca Vadalà, MD, PhD<sup>1,2</sup>, Amit Jain, MD, MBA<sup>10</sup> ,  
Andreas K. Demetriades, MD<sup>11</sup> , Sam K. Cho, MD<sup>12</sup> , Patrick C. Hsieh, MD<sup>13</sup> ,  
Ashish D. Diwan, MD, PhD<sup>14,15</sup>, Christopher Martin, MD<sup>16</sup>, Tim Yoon, MD, PhD<sup>17</sup> ,  
Sathish Muthu, MS, PhD<sup>18,19,20</sup> , and AO Spine Knowledge Forum Degenerative

## Abstract

**Study Design:** Cross-sectional study.

**Objectives:** To assess global practices and preferences in the use of osteobiologics for anterior cervical discectomy and fusion (ACDF) and identify factors influencing the choice of specific osteobiologics.

**Methods:** An online survey developed by AO Spine was distributed to spine surgeons worldwide. The survey captured demographic characteristics, osteobiologic use and related information (i.e., previous training, practice patterns, etc.), and

<sup>1</sup> Operative Research Unit of Orthopaedic and Trauma Surgery, Fondazione Policlinico Universitario Campus Bio-Medico, Rome, Italy

<sup>2</sup> Research Unit of Orthopaedic and Trauma Surgery, Department of Medicine and Surgery, Università Campus Bio-Medico di Roma, Rome, Italy

<sup>3</sup> Reva Spine Centre, Visakhapatnam, India

<sup>4</sup> Department of Orthopaedics, Noordwest Hospitals, Alkmaar, The Netherlands

<sup>5</sup> Gerling Institute, New York City, NY, USA

<sup>6</sup> Department of Neurosurgery, BG Klinikum Bergmannstrost Halle, Halle, Germany

<sup>7</sup> Department of Health Research Methods Evidence and Impact, McMaster University, Hamilton, ON, Canada

<sup>8</sup> Department of Orthopaedics and Traumatology, The University of Hong Kong, Hong Kong SAR, China

<sup>9</sup> Department of Orthopaedic Surgery, University of California, Davis, CA, USA

<sup>10</sup> Department of Orthopaedic Surgery, Johns Hopkins University, Baltimore, MD, USA

<sup>11</sup> Department of Neurosurgery, Edinburgh Spinal Surgery Outcome Studies Group, Royal Infirmary Edinburgh, Edinburgh, UK

<sup>12</sup> Department of Orthopedic Surgery, Icahn School of Medicine at Mount Sinai, New York, NY, USA

<sup>13</sup> Department of Neurological Surgery, Keck School of Medicine, University of Southern California, Sacramento, CA, USA

<sup>14</sup> Spine Labs, St George and Sutherland Clinical School, University of New South Wales, Kogarah, NSW, Australia

<sup>15</sup> Spine Service, Department of Orthopaedic Surgery, St George and Sutherland Clinical School, University of New South Wales, Kogarah, NSW, Australia

<sup>16</sup> Department of Orthopaedic Surgery, University of Minnesota, Minneapolis, MN, USA

<sup>17</sup> Department of Orthopedics, Emory University, Atlanta, GA, USA

<sup>18</sup> Department of Orthopedic Surgery, Orthopedic Research Group, Coimbatore, India

<sup>19</sup> Department of Biotechnology, Faculty of Engineering, Karpagam Academy of Higher Education, Coimbatore, India

<sup>20</sup> Department of Orthopaedics, Government Medical College, Karur, India

## Corresponding Author:

Sathish Muthu, Department of Orthopedics, Government Hospital, Velayuthampalayam, Karur 639117, India.

Email: [drsathishmuthu@gmail.com](mailto:drsathishmuthu@gmail.com)



Creative Commons Non Commercial No Derivs CC BY-NC-ND: This article is distributed under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs 4.0 License (<https://creativecommons.org/licenses/by-nc-nd/4.0/>) which permits non-commercial use, reproduction and distribution of the work as published without adaptation or alteration, without further permission provided the original work is attributed as specified on the SAGE and Open Access pages (<https://us.sagepub.com/en-us/nam/open-access-at-sage>).

factors influencing osteobiologic choice in ACDF. Descriptive statistics, Chi-square tests, and multiple logistic regression were used to analyze responses, focusing on the associations between osteobiologic use and variables such as training, cost awareness, and regional practices.

**Results:** Responses from 458 surgeons revealed regional variability in osteobiologic preferences. Autologous iliac crest bone graft (AICBG) was predominant in Asia Pacific and Middle East, while allograft and demineralized bone matrix were favored in North America and Latin America ( $P < 0.0001$ ). Over half of the respondents (79.7%) lacked formal training in osteobiologics, and 53.1% were unaware of related costs. Surgeons residing in the Asia Pacific region (OR: 0.47, 95% CI: 0.26-0.84,  $P = 0.0114$ ), without formal training (OR: 0.53, 95% CI: 0.29-0.97,  $P = 0.0429$ ), or using cages less often (OR: 0.15, 95% CI: 0.06-0.34,  $P < 0.0001$ ) were less likely to utilize osteobiologics. Osteobiologic use was also more common when related costs were not an issue for the practitioner (OR: 2.32, 95% CI: 1.47-3.70,  $P = 0.0004$ ).

**Conclusions:** Significant variation exists in osteobiologic use in ACDF across global regions, influenced by surgeon training, cost awareness, and institutional resources. Enhanced training and guidelines could improve consistency in osteobiologic application.

## Keywords

anterior cervical discectomy and fusion, ACDF, osteobiologic, bone graft, allograft, cervical spine, autograft, fusion

## Introduction

Anterior cervical discectomy and fusion (ACDF) is one of the most commonly performed spine surgeries worldwide, with a high success rate ranging from 85 to 95%.<sup>1</sup> The primary objectives of ACDF are to relieve pain and restore neurological function, which are achieved through thorough decompression of neural structures. However, long-term success largely depends on achieving solid bony fusion.<sup>2</sup> This can be attained through various combinations of implants (e.g., cages and plates) and osteobiologics.

Osteobiologics are substances that can induce bone formation (osteoinduction), support bone ingrowth (osteocondution), or directly generate new bone tissue (osteogenesis).<sup>3</sup> Autologous iliac crest bone graft (AICBG) has traditionally been considered the gold standard for achieving fusion due to its ready availability and excellent osteogenic, osteoinductive, and osteoconductive properties. Despite these advantages, AICBG is associated with significant donor site morbidity, including wound complications, persistent pain, infection, and hematoma.<sup>4</sup>

The limitations of AICBG have prompted the development of a variety of osteobiologics to augment or replace autografts. These include allografts, bone morphogenetic protein (BMP)-2, demineralized bone matrix (DBM), hydroxyapatite (HA), beta tricalcium phosphate ( $\beta$ -TCP), bioglass, bioceramics, and synthetic peptides. While these materials offer potential advantages, their use is complicated by variability in clinical evidence supporting their efficacy and safety. Indeed, there is a lack of high-quality clinical evidence for the use of osteobiologics, which provides limited information to foster the knowledge and to support their use.<sup>5-7</sup> Furthermore, there is ongoing debate whether the use of osteobiologics, either AICBG or other grafts, significantly affects fusion rates in revision ACDF cases, where the risk of pseudoarthrosis is

substantially higher.<sup>8</sup> Additionally, regulatory differences across countries, including variations in reimbursement policies and market availability, contribute to inconsistent adoption among spine surgeons.<sup>9,10</sup> From a global market perspective, this is particularly relevant since the economic potential of osteobiologics has been estimated >\$9 billion in 2024.<sup>11</sup> These disparities highlight a critical need to standardize practices and evaluate the factors influencing clinical decision-making.

Given the limited clinical data available, there is no consensus among spine surgeons regarding the use of appropriate osteobiologics in specific clinical scenarios, whether for single- or multi-level ACDF. AO Spine recently developed an international guideline, the *AO Spine Guideline for the Use of Osteobiologics (AOGO)*, to provide evidence-based recommendations.<sup>12</sup> The aim of the current study is to assess global practice patterns and preferences regarding the use of osteobiologics in ACDF, and to identify the factors influencing the selection of specific osteobiologics. This research addresses a critical gap by exploring how individual surgeon preferences, familiarity, and access to resources drive decision-making. Additionally, it seeks to evaluate whether cost-effectiveness and clinical evidence play a secondary role in the selection process. By elucidating these dynamics, this study aims to inform future research, enhance the applicability of guidelines, and promote consensus among spine surgeons.

## Materials and Methods

### Study Participants

An online questionnaire assessing the use of osteobiologics in ACDF was formulated by the AO Spine Knowledge Forum Degenerative working group within the AOGO project. AO Spine is a professional medical association of spine surgeons and a clinical division of the AO Foundation. AO Spine is the

leading global academic community of spine surgeons dedicated to generating, distributing, and sharing knowledge through research, education and community development. Established in June 2003, the organization is led by the AO Spine International Board. The board guides and supports the AO Spine regions Asia Pacific, Europe and Africa, Latin America, Middle East, and North America. AO Spine fosters a global community of 30,000 members and associates from all over the world. Within the society, the Knowledge Forum Degenerative is one of five expert-driven working groups generating evidence-based knowledge on degenerative disorders of the spine. The survey was developed within the AO Spine Knowledge Forum Degenerative via iterative review cycles among the members of the group which are key opinion leaders in the field in the Degenerative Spine surgery field. No formal institutional review board approval was required for this study. The survey was sent out by email to approximately 6,000 spine surgeons between March 1 and 15, 2024. The link was kept accessible for 15 days with two reminders sent out during the period. All the participants signed a digital informed consent and agreed on the use of their anonymized responses for research purposes.

### Study Questionnaire

The survey originally included 23 questions. General participants' demographics included country and AO Spine region of practice, gender, age, years of practice in spine surgery, specialty, practice setting, information about spine surgery fellowship, number of spine surgery cases performed per year, and number of ACDF surgeries performed per year. Then, information regarding previous formal training in the use of osteobiologics, awareness of osteobiologic costs, use of implants in ACDF surgeries (i.e., cages and plates), frequency of osteobiologic use in ACDF, strategies to reduce complications and improve treatment outcomes, and factors influencing the use of osteobiologics in ACDF. These factors included cost, evidence base, historical practice, availability, education, and vendor support, and were rated by each participant from 1 ("most important") to 6 ("not important"). The full questionnaire is available as a [Supplemental Material](#).

### Statistical Analysis

Categorical data were shown as absolute (n) and relative (%) frequencies. Statistical analysis of data was performed using the Chi-square test. Multiple logistic regression was performed to investigate the associations between the use of osteobiologics, related training, and different instrumentation with AO Spine region, hospital setting, community setting, surgeon's age, years of surgical practice, overall spine surgery and ACDF case volume, specialty, fellowship, knowledge of osteobiologic costs, existence of institutional cost limitations, and use of local corticosteroids. These covariates were

selected based on their statistical significance as per univariate analyses ([Supplemental Tables](#)). Answers reporting the use of multiple osteobiologics were removed from the analysis. Reference levels were selected based on their frequency or their relevance per each independent variable. Odds ratios (OR) and 95% confidence intervals (CI) were estimated for each reference category. Statistical significance was set at  $P < 0.05$ . Formal analysis was performed using Prism 10 (v. 10.1.1, GraphPad Software).

## Results

### Participants' Demographics

A total of 458 surgeons completed the survey. Participants were from North America (12.2%), Latin America (14.4%), Europe & Southern Africa (35.2%), Middle East & North Africa (13.5%), and Asia Pacific (24.7%) regions. The majority of respondents were male (94.6%), aged between 35 and 44 (39.1%), and had been practicing for at least 5 years (79.7%). Among these, 57.6% were orthopaedic surgeons, 40.4% practiced in an academic hospital, 28.4% in a private setting, and 27.5% in a public or government/military hospital; most worked in urban areas (79.7%). The majority had completed a spine surgery fellowship (65.1%): of these, one-third finished their post-graduate training within the last 5 years, one-fourth within the last 5 to 10 years, and the rest more than 10 years ago. Regarding surgical volume, 26.9% performed fewer than 100 spine surgeries per year, 36.2% operated between 100 and 200 cases annually, and the remaining surgeons operated on more than 200 cases per year. Among these, 45.0% performed fewer than 20 yearly ACDFs, 36.7% performed between 20 and 50, and the rest performed over 50 ACDFs annually.

### Use of Implants and Osteobiologics during ACDF Surgery

Among all participants, 79.7% reported not having received any formal training in the use of osteobiologics. Of those who did receive training, it was mainly through courses (10.5%, including events hosted by AO Spine, North American Spine Society [NASS], Cervical Spine Research Society, and American Associations of Neurological Surgeons), webinars (6.8%), individual training (4.4%), educational events organized by the industry (2.8%), hands-on experience (2.0%), and fellowship training (1.7%). A slight majority of respondents (53.1%) were not aware of the costs associated with osteobiologic use at their institution. According to univariate analyses, osteobiologic-related training ([Table 1](#)) was significantly different when considering the regional distribution of participants ( $P < 0.0001$ ), years in practice ( $P = 0.0274$ ), fellowship training ( $P = 0.0023$ ), knowledge of osteobiologic costs ( $P = 0.0004$ ), use of plates ( $P = 0.0085$ ), and corticosteroids ( $P = 0.0222$ ).

**Table 1.** Univariate Analyses Assessing the Relationship Between Osteobiologic-related Training and Study Covariates.

Covariate	Trained	Not trained	P
AO Spine region			<b>&lt;0.0001</b>
Europe and Southern Africa	26	135	
Asia Pacific	22	91	
Latin America	15	51	
North America	24	32	
Middle East and Northern Africa	6	56	
Age group			0.3034
25-34	6	44	
35-44	32	147	
45-54	33	102	
55-64	14	48	
65+	8	24	
Years in practice			<b>0.0274</b>
<5	17	76	
5-10	22	93	
11-15	10	69	
16-20	20	37	
>20	24	90	
Specialty			0.2017
Orthopaedic surgeon	59	204	
Neurosurgeon	31	155	
Other	3	6	
Practice setting			0.0755
Private practice	22	108	
Academic/University affiliate	47	138	
Public hospital or Government/ Military hospital	19	107	
Other	5	12	
Community			0.4529
Urban	75	290	
Suburban	17	62	
Rural	1	13	
Fellowship training			<b>0.0023</b>
Yes	73	225	
No	20	140	
Spine surgical volume (cases/year)			0.4959
<100	21	102	
101-200	33	133	
201-300	24	80	
301-400	10	24	
>400	5	26	
ACDF surgical volume (cases/year)			0.2406
1-20	35	171	
20-50	37	131	
>50	21	63	
Informed about osteobiologic cost			<b>0.0004</b>
Yes	59	156	
No	34	209	

**Table 1.** (continued)

Covariate	Trained	Not trained	P
Use of interbody cages			0.1317
Yes	80	333	
No	13	32	
Use of plate			<b>0.0085</b>
Yes	65	200	
No	28	165	
Use of local CCS			<b>0.0222</b>
Yes	23	54	
No	70	311	
Cost limitation			0.4393
Yes	34	118	
No	59	247	

Statistically significant P values are shown in bold.

During ACDF surgery, most surgeons (68.1%) reported always using interbody cages. Not using cages was generally uncommon with other respondents, who employed these constructs most of the time (16.8%); around 50% of the time (5.2%); rarely (5.5%); or never (4.4%). Univariate analyses showed that the use of cages (Table 2) significantly differed based on regionality ( $P = 0.0007$ ), age ( $P = 0.0025$ ), years in practice ( $P = 0.0026$ ), specialty ( $P = 0.0210$ ), spine surgical volume ( $P = 0.0216$ ), ACDF case volume ( $P = 0.0493$ ), knowledge of osteobiologic costs ( $P = 0.0106$ ), and anterior plating ( $P = 0.0269$ ).

Slightly more than one-third of respondents rarely used anterior plating (34.7%), whereas 26.0% always used it, 17.7% used it most of the time, 14.2% used it around 50.0% of the time, and 7.4% never used it. The use of anterior plating (Table 3) was significantly different when considering regional distribution ( $P < 0.0001$ ), specialty ( $P < 0.0001$ ), and use of interbody cages ( $P = 0.0269$ ).

In terms of using osteobiologics as a supplement to autologous local bone, 38.0% of the surgeons always used some form of osteobiologic, 17.3% used them most of the time, 7.9% used them half of the time, while 22.5% and 14.4% used them rarely or never, respectively. As an alternative to autologous local bone graft, 28.0% reported using a tricortical bone graft, 27.3% used DBM, 22.7% used allograft, 12.7% used HA, 9.4% used cancellous AICBG, 7.7% used cellular bone graft, 7.4% used bicalcium phosphate, 6.6% used BMP-2, 1.3% used ABM/P-15 peptide, and a small percentage used other grafts including tibial (0.2%) and fibular grafts (0.2%), unspecified synthetic materials (0.2%), bone marrow aspirate (0.2%),  $\beta$ -TCP (0.2%), collagen matrix (0.2%), bioactive glass (0.8%), or no graft at all (1.0%). When subgrouping the three most common osteobiologics by regions, AICBG was the most popular in Asia Pacific (58.5%) and Middle East and Northern Africa (45.3%), allograft was primarily employed in

(continued)

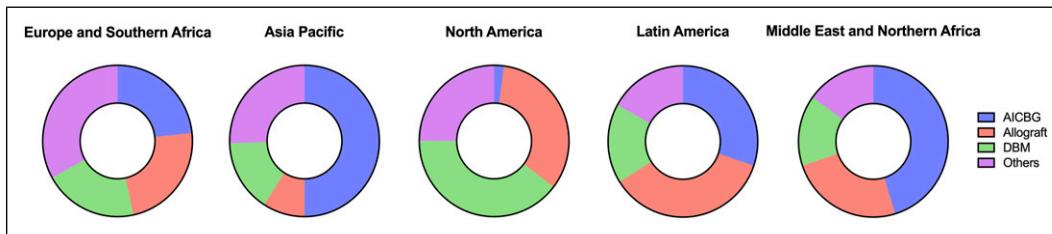
**Table 2.** Univariate Analyses Assessing the Relationship Between the Use of Interbody Cages (>50% of Cases) and Study Covariates.

Covariate	Cage	No Cage	<i>p</i>
AO Spine region			<b>0.0007</b>
Europe and Southern Africa	149	12	
Asia Pacific	104	9	
Latin America	65	1	
North America	46	10	
Middle East and Northern Africa	49	13	
Age group			<b>0.0025</b>
25-34	39	11	
35-44	172	7	
45-54	119	16	
55-64	55	7	
65+	28	4	
Years in practice			<b>0.0026</b>
<5	74	19	
5-10	107	8	
11-15	76	3	
16-20	52	5	
>20	104	10	
Specialty			<b>0.0210</b>
Orthopaedic surgeon	234	29	
Neurosurgeon	173	13	
Other	6	3	
Practice setting			0.2151
Private practice	119	11	
Academic/University affiliate	151	24	
Public hospital or Government/Military hospital	117	9	
Other	16	1	
Community			0.4529
Urban	75	290	
Suburban	17	62	
Rural	1	13	
Fellowship training			0.0821
Yes	274	24	
No	139	21	
Spine surgical volume (cases/year)			<b>0.0216</b>
<100	102	21	
101-200	131	15	
201-300	100	4	
301-400	32	2	
>400	28	4	
ACDF surgical volume (cases/year)			<b>0.0493</b>
1-20	178	28	
20-50	157	11	
>50	78	6	
Informed about osteobiologic cost			<b>0.0106</b>
Yes	202	13	
No	211	32	
Use of plate			<b>0.0269</b>
Yes	232	33	
No	181	12	
Use of local CCS			0.0563
Yes	70	7	
No	343	38	
Cost limitation			0.0913
Yes	132	20	
No	281	25	

Statistically significant *P* values are shown in bold.**Table 3.** Univariate Analyses Assessing the Relationship Between the Use of an Anterior Plate (>50% of Cases) and Study Covariates.

Covariate	Plate	No Plate	<i>p</i>
AO Spine region			<b>&lt;0.0001</b>
Europe and Southern Africa	63	98	
Asia Pacific	77	36	
Latin America	35	31	
North America	52	4	
Middle East and Northern Africa	38	24	
Age group			0.4038
25-34	30	20	
35-44	112	67	
45-54	70	65	
55-64	36	26	
65+	17	15	
Years in practice			0.0510
<5	64	29	
5-10	71	44	
11-15	39	40	
16-20	32	25	
>20	59	55	
Specialty			<b>&lt;0.0001</b>
Orthopaedic surgeon	176	87	
Neurosurgeon	83	103	
Other	6	3	
Practice setting			0.3582
Private practice	69	61	
Academic/University affiliate	115	70	
Public hospital or Government/Military hospital	70	56	
Other	11	6	
Community			0.2379
Urban	204	161	
Suburban	52	27	
Rural	9	5	
Fellowship training			0.9330
Yes	172	126	
No	93	67	
Spine surgical volume (cases/year)			0.4491
<100	76	47	
101-200	98	68	
201-300	58	46	
301-400	15	19	
>400	18	13	
ACDF surgical volume (cases/year)			0.4450
1-20	125	81	
20-50	91	77	
>50	49	35	
Informed about osteobiologic cost			0.4948
Yes	128	87	
No	137	106	
Use of interbody cages			<b>0.0269</b>
Yes	232	181	
No	33	12	
Use of local CCS			0.1028
Yes	51	26	
No	214	167	
Cost limitation			0.2239
Yes	94	58	
No	171	135	

Statistically significant *P* values are shown in bold.



**Figure 1.** Regional distribution of the three most common osteobiologics. Abbreviations: AICBG = autologous iliac crest bone graft; DBM = demineralized bone matrix.

Latin America (35.8%) and North America (33.3%), and DBM was mostly used in North America (39.6%) and Europe and Southern Africa (20.3%). All these differences were statistically significant ( $P < 0.0001$ ; **Figure 1**). The use of osteobiologics (**Table 4**) showed statistically significant differences when considering geographical distribution ( $P = 0.0006$ ), osteobiologic-related training ( $P = 0.0453$ ), specialty ( $P = 0.0178$ ), community ( $P = 0.0271$ ), fellowship training ( $P = 0.0317$ ), ACDF case volume ( $P = 0.0305$ ), knowledge of osteobiologic costs ( $P = 0.0279$ ), use of interbody cages ( $P < 0.0001$ ), and limitations related to osteobiologic use ( $P < 0.0001$ ).

Regarding strategies to reduce the risk of complications, 16.8% of surgeons routinely used topical corticosteroids to reduce the risk of dysphagia, and 16.4% described various specific strategies, including applied substances or technical tricks, to improve outcomes and/or reduce complications. These strategies have been summarized in **Table 5**.

When asked about the main factors influencing their choice of a specific osteobiologic, the most important was the evidence base, followed by availability, cost, historical practice, education, and vendor support (**Figure 2**). In 66.8% of cases, respondents were not limited by the costs of osteobiologics. However, most of the remaining surgeons (23.6%) were burdened by the high costs of these products, especially when practicing in low-to-middle-income countries and under healthcare systems in which insurances do not cover for osteobiologics, thus requiring patients to pay out of their own pockets. Additional factors limiting the utilization of these products were the lack of strict necessity for their use given acceptable outcomes with other approaches (3.5%), local hospital policies (3.9%), and product availability (1.3%). Finally, 91.5% of surgeons were not aware of specific guidelines regarding the use of osteobiologics in ACDF. Conversely, the remaining surgeons mentioned guidelines from AO Spine (5.0%), NASS (0.7%), or other sources (2.8%).

### Factors Associated with the Use of Osteobiologics

The use of osteobiologics to augment autologous local bone (**Table 6**) was significantly less popular among participants from the Asia Pacific region (OR: 0.47, 95% CI: 0.26-0.84,  $P = 0.0114$ ), those who did not receive

formal training (OR: 0.53, 95% CI: 0.29-0.97,  $P = 0.0429$ ), and in those who used a cage in <50% of cases (OR: 0.15, 95% CI: 0.06-0.34,  $P < 0.0001$ ). Conversely, osteobiologic use was significantly more common when related costs were not an issue for the practitioner (OR: 2.32, 95% CI: 1.47-3.70,  $P = 0.0004$ ). Furthermore, these participants showed lower odds of using both a tricortical bone graft (OR: 0.23, 95% CI: 0.11-0.45,  $P < 0.0001$ ) and an iliac crest cancellous bone graft (OR: 0.25, 95% CI: 0.10-0.66,  $P = 0.054$ ).

Formal training in the use of osteobiologics (**Table 7**) was significantly more common among surgeons from North America (OR: 2.30, 95% CI 1.05-5.05,  $P = 0.0377$ ) and in practice for >20 years (OR: 2.34, 95% CI 1.04-5.35,  $P = 0.0417$ ). Conversely, osteobiologic-related training was less popular among participants who did not do a fellowship (OR: 0.53, 95% CI: 0.29-0.94,  $P = 0.0327$ ) and not informed about osteobiologic costs (OR: 0.49, 95% CI: 0.29-0.80,  $P = 0.0049$ ).

The use of a cage in >50% of ACDF cases (**Table 8**) was significantly less common in surgeons aged more than 45 years (45-54: OR: 0.08, 95% CI 0.01-0.71,  $P = 0.0272$ ; 55-64: OR: 0.16, 95% CI 0.03-0.77,  $P = 0.0269$ ; 65+: OR: 0.07, 95% CI 0.01-0.83,  $P = 0.0373$ ) compared to younger peers. However, interbody cages were also increasingly utilized in surgeons with more than 10 years of practice (11-15: OR: 21.04, 95% CI 3.68-159-40,  $P = 0.0013$ ; 16-20: OR: 17.80, 95% CI 22.60-151.70,  $P = 0.0052$ ; >20: OR: 13.87, 95% CI 2.46-94.26,  $P = 0.0044$ ). In addition, the use of a cage was significantly more common when anterior plating was not employed (OR: 3.08, 95% CI: 1.29-7.79,  $P = 0.0138$ ).

The use of a plate in >50% of ACDF cases (**Table 9**) was significantly more common in Asia Pacific (OR: 2.63, 95% CI: 1.49-4.70,  $P = 0.0009$ ), Middle East and Northern Africa (OR: 2.18, 95% CI: 1.09-4.36,  $P = 0.0272$ ), and North America (OR: 25.02, 95% CI 9.31-87.70,  $P < 0.0001$ ).

### Discussion

The findings of this study reveal important insights into the global practices and preferences for osteobiologics in ACDF surgery. The high variability in the use of osteobiologics,

**Table 4.** Univariate Analyses Assessing the Relationship Between Osteobiologic Use and Study Covariates.

Covariate	Used	Not Used	p
AO Spine region			<b>0.0006</b>
Europe and Southern Africa	109	52	
Asia Pacific	58	55	
Latin America	49	17	
North America	31	31	
Middle East and Northern Africa	42	14	
Osteobiologic-related training			<b>0.0453</b>
Yes	67	26	
No	222	143	
Age group			0.2453
25-34	27	23	
35-44	111	68	
45-54	86	47	
55-64	33	27	
65+	19	13	
Years in practice			0.5470
<5	55	38	
5-10	70	45	
11-15	52	27	
16-20	41	16	
>20	71	43	
Specialty			<b>0.0178</b>
Orthopaedic surgeon	162	101	
Neurosurgeon	125	61	
Other	2	7	
Practice setting			0.8957
Private practice	84	46	
Academic/University affiliate	118	67	
Public hospital or Government/Military hospital	76	50	
Other	11	6	
Community			<b>0.0271</b>
Urban	232	133	
Suburban	52	27	
Rural	5	11	
Fellowship training			<b>0.0317</b>
Yes	194	99	
No	89	70	
Spine surgical volume (cases/year)			0.1377
<100	73	50	
101-200	110	56	
201-300	65	39	
301-400	26	8	
>400	15	16	
ACDF surgical volume (cases/year)			<b>0.0305</b>
1-20	101	85	
20-50	111	57	
>50	57	27	
Informed about osteobiologic cost			<b>0.0279</b>
Yes	147	68	
No	142	101	
Use of interbody cages			<b>&lt;0.0001</b>
Yes	278	135	
No	11	34	
Use of plate			0.6307
Yes	164	101	
No	125	68	
Use of local CCS			0.2532
Yes	53	24	
No	236	145	
Cost limitation			<b>&lt;0.0001</b>
Yes	73	79	
No	216	90	

Statistically significant P values are shown in bold.

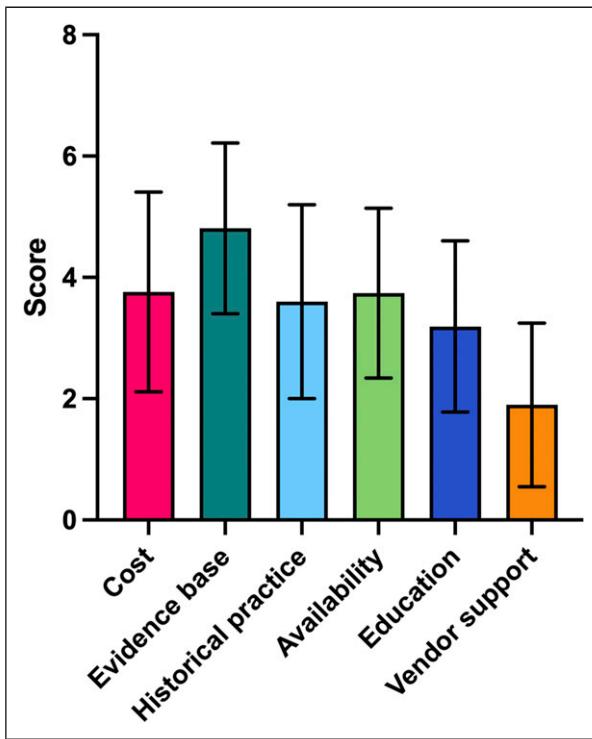
**Table 5.** Strategies and Tricks to Avoid Complications as Reported by the Surveyed Participants.

Employed Strategy	N (%)
Use or non-fixed retractors	6 (1.3)
Intermittent retractor release	2 (0.4)
Frequent inflation and deflation of the endotracheal cuff	6 (1.3)
Application of reduced cuff pressure	4 (0.8)
Application of vancomycin powder or other antibiotics or antiseptics (e.g., iodopovidone)	12 (2.6)
Placement of gel foam soaked in dexamethasone overlying the plate	1 (0.2)
Use of nasogastric tube	1 (0.2)
Interbody placement of gel foam soaked in the patient's blood and mixed with local bone	1 (0.2)
Use of local anesthesia	3 (0.6)
Application of local steroids	3 (0.6)
Controlled tracheoesophageal traction	1 (0.2)
Placement of a cellulose pad behind the esophagus	1 (0.2)
Solid meals by mouth postponed 24 h after surgery	1 (0.2)
Copious irrigation	2 (0.4)
Use of topical hemostatic agents to control bleeding (e.g., Surgiflo®, tranexamic acid, gel foam, etc.)	14 (3.1)
Inhalatory corticosteroids for 72 h and mannitol for 24 h before surgery	1 (0.2)
Administration of systemic steroids	2 (0.4)
Hole in the endplate to allow for bleeding	2 (0.4)
Preoperative tracheal traction exercises	1 (0.2)
Drain placement	5 (1.1)
Prescription of postoperative orthosis	1 (0.2)
Local cooling	1 (0.2)
No drilling	1 (0.2)

both in terms of types and frequency, underscores the lack of a universally accepted standard in clinical practice. This variability is influenced by multiple factors including regional differences, surgeon training, and economic considerations.

One of the main observations is the significant proportion of surgeons who have not received formal training in the use of osteobiologics. This highlights a critical gap in spine surgery education, suggesting that despite the availability of osteobiologics and their potential benefits, many surgeons may be underprepared to effectively utilize these materials. Given that the effectiveness and safety of each osteobiologic are closely linked to its underlying mechanisms and specific biological pathways, a deep understanding of how these materials function *in vivo* is essential.<sup>13</sup> Overlooking these nuances may potentially cause undesired effects and severe adverse events, which might even be life-threatening.<sup>14</sup> Furthermore, whether osteobiologics significantly improve ACDF clinical outcomes and fusion rates is still to be determined, as recently highlighted by a systematic review.<sup>15</sup>

The reliance on unregulated learning platforms such as courses, webinars, and industry-sponsored events may



**Figure 2.** Main factors influencing the choice of a specific osteobiologic among participants rated between 1 (“most important”) and 6 (“not important”).

contribute to the inconsistent use of osteobiologics, as these methods do not always provide the comprehensive training necessary for informed clinical decision-making. Furthermore, over relying on industry-organized educational events may skew perceptions of efficacy towards the products being promoted, increasing the risk of conflicts of interest.

The study also reveals a lack of awareness regarding the costs of osteobiologics among many surgeons, which could influence their adoption and utilization. This is particularly relevant considering that the costs associated with ACDF widely vary from hundreds to several thousand dollars based on the specific implants and osteobiologics employed,<sup>16</sup> as well as due to local reimbursement policies and insurance coverage.<sup>17</sup> In regions where healthcare systems do not cover the cost of these materials, surgeons may be hesitant to use them, particularly if patients must bear the financial burden.<sup>18</sup> This economic barrier could lead to disparities in the quality of care provided, with patients in lower-income regions potentially receiving less optimal or more invasive treatments (i.e., due to the higher use of AICBG) because of cost constraints.

Another critical factor influencing osteobiologic use is the evidence base supporting their efficacy. Surgeons were more likely to adopt osteobiologics when there is strong, evidence-based support for their use. However, this study shows that in the absence of robust evidence, historical

**Table 6.** Multivariate Logistic Regression Model Showing the Association Between Osteobiologic Use in ACDF and Study Covariates.

Covariate	OR	95% CI	p
<b>AO Spine Region</b>			
Europe and Southern Africa (reference)			
Asia Pacific	0.47	0.26-0.84	<b>0.0114</b>
Latin America	1.22	0.62-2.50	0.5761
North America	1.47	0.66- 3.47	0.3612
Middle East and Northern Africa	0.55	0.27-1.12	0.0987
<b>No formal osteobiologic-related training</b>	0.53	0.29- 0.97	<b>0.0429</b>
<b>Specialty</b>			
Orthopaedic surgeon (reference)			
Neurosurgeon	1.43	0.87- 2.34	0.1626
Other	0.22	0.03-1.21	0.0968
<b>Community</b>			
Urban (reference)			
Suburban	1.51	0.82- 2.83	0.1935
Rural	0.22	0.03-0.96	0.0692
<b>No fellowship training</b>	0.67	0.40-1.13	0.1308
<b>ACDF surgical volume (cases/year)</b>			
1-20 (reference)			
20-50	1.07	0.63-1.80	0.7986
50-100	1.33	0.66-2.70	0.4285
>100	0.86	0.27-2.94	0.8077
<b>Not informed about osteobiologic-related costs</b>	0.87	0.56-1.37	0.5566
<b>No use of interbody cages</b>	0.15	0.06-0.34	<b>&lt;0.0001</b>
<b>No cost limitations on osteobiologics</b>	2.32	1.47-3.70	<b>0.0004</b>
<b>Type of osteobiologic</b>			
Tricortical bone graft	0.23	0.11-0.45	<b>&lt;0.0001</b>
Iliac crest cancellous bone graft	0.25	0.10-0.66	<b>0.0054</b>

Abbreviations: ACDF = anterior cervical discectomy and fusion, CI = confidence interval, OR = odds ratio. Statistically significant P values are shown in bold

practices, and surgeon familiarity often guide clinical decisions. This also applies to some of technical tricks employed by surveyed participants, such as the use of inhalatory corticosteroids, retroesophageal padding, or the use of vancomycin powder, which has not been validated in ACDF neither proven definitively safe and effective in other spine regions.<sup>19,20</sup> This reliance on tradition over evidence can perpetuate the use of less effective, outdated, or even harmful techniques, further contributing to the variability in ACDF outcomes.

The regional differences in osteobiologic use observed in this study also raise important considerations. For instance, surgeons in the Asia Pacific region were less likely to utilize osteobiologics, and reported a substantially more frequent use of AICBG, either in the form of a tricortical or cancellous bone graft. Conversely, osteobiologic-related training was

**Table 7.** Multivariate Logistic Regression Model Showing the Association Between Osteobiologic-related Training and Study Covariates.

Covariate	OR	95% CI	p
<b>AO Spine Region</b>			
Europe and Southern Africa (reference)			
Asia Pacific	0.99	0.50-1.95	0.9757
Latin America	1.35	0.63-2.82	0.4282
North America	2.30	1.05-5.05	<b>0.0377</b>
Middle East and Northern Africa	0.58	0.20-1.47	0.2751
<b>No fellowship training</b>	0.53	0.29-0.94	<b>0.0327</b>
<b>Years in practice</b>			
<5 (reference)			
5-10	1.14	0.54-2.44	0.7344
11-15	0.63	0.25-1.52	0.3082
16-20	1.02	0.48-2.18	0.9654
>20	2.34	1.04-5.35	<b>0.0417</b>
<b>Not informed about osteobiologic-related costs</b>	0.49	0.29-0.80	<b>0.0049</b>
<b>No use of plate</b>	0.81	0.38-1.81	0.6015
<b>Use of local CCS</b>	1.43	0.76-2.59	0.2535

Abbreviations: CCS = corticosteroids, CI = confidence interval, OR = odds ratio. Statistically significant P values are shown in bold.

significantly more common in North American surgeons, which further highlights regional discrepancies in training pathways. These variable practice patterns may be due to differences in access to certain medical technologies, variations in healthcare infrastructures, or cultural and educational differences in surgical training. Such regional discrepancies highlight the need for more standardized global guidelines and improved access to training and resources.<sup>21</sup>

The lack of awareness and use of guidelines related to the use of osteobiologics in ACDF is another concerning finding. Indeed, 9 out of 10 participants were unaware of the existence of any specific documentation, and the remaining sparingly mentioned guidelines from NASS and AO Spine. Over the years, NASS has issued evidence-based coverage recommendations for various materials, including allograft and DBM (2017), BMP-2 (2014), and synthetics (2023).<sup>22</sup> However, a comprehensive guideline has yet to be released. In this context, the AOGO guideline marks a significant milestone being the first to offer recommendations for the use of osteobiologics in ACDF.<sup>12</sup> Interestingly, the results of the present survey align closely with AOGO recommendations. The widespread use of structural AICBGs or cages with cancellous AICBGs or other osteobiologics such as allograft, DBM, HA, and  $\beta$ -TCP reflects the guidance advocating for either autograft or allograft vs. cages with osteobiologics, both of which are associated with comparable outcomes.<sup>12</sup> However, the low awareness of these guidelines suggests that more efforts are needed to disseminate and implement them across the global spine surgery community. Given that these

**Table 8.** Multivariate Logistic Regression Model Showing the Association Between the Use of Interbody Cages (>50% of Cases) and Study Covariates.

Covariate	OR	95% CI	p
<b>AO Spine Region</b>			
Europe and Southern Africa (reference)			
Asia Pacific	0.52	0.16-1.66	0.2671
Latin America	6.14	0.99-120.40	0.1031
North America	0.85	0.27-2.73	0.7784
Middle East and Northern Africa	0.51	0.16-1.66	0.0700
<b>Age</b>			
25-34 (reference)			
35-44	3.55	1.00-13.31	0.0524
45-54	0.08	0.01-0.71	<b>0.0272</b>
55-64	0.16	0.03-0.77	<b>0.0269</b>
65+	0.07	0.01-0.83	<b>0.0373</b>
<b>Years in practice</b>			
<5 (reference)			
5-10	3.54	1.01-14.09	0.0569
11-15	21.04	3.68-159.40	<b>0.0013</b>
16-20	17.80	2.60-151.70	<b>0.0052</b>
>20	13.87	2.46-94.26	<b>0.0044</b>
<b>Specialty</b>			
Orthopaedic surgeon (reference)			
Neurosurgeon	1.08	0.48-2.51	0.8531
Other	0.13	0.01-2.99	0.1076
<b>Spine surgical volume (cases/year)</b>			
<100 (reference)			
100-200	1.29	0.48-3.47	0.6150
201-300	2.20	0.51-10.77	0.3034
301-400	1.50	0.24-13.40	0.6843
>400	0.81	0.14-5.64	0.8168
<b>ACDF surgical volume (cases/year)</b>			
1-20 (reference)			
20-50	2.79	1.02-8.20	0.0525
50-100	2.37	0.62-10.20	0.2229
>100	2.96	0.29-76.04	0.4154
<b>Not informed about osteobiologic-related costs</b>	0.55	0.24-1.22	0.1499
<b>No use of plate</b>	3.08	1.29-7.79	<b>0.0138</b>

Abbreviations: ACDF = anterior cervical discectomy and fusion, CI = confidence interval, OR = odds ratio. Statistically significant P values are shown in bold.

guidelines were only recently introduced, it is likely that they will reach a broader audience over time, leading to greater implementation of the recommended practices in the near future. Therefore, the authors plan to repeat the survey after a designated period to evaluate the extent of guideline adoption and track changes in practice over time following their implementation.

This study has some limitations. First, the overall low response rate (458 respondents out of approximately 6,000 recipients) may have limited the reliability of the collected

**Table 9.** Multivariate Logistic Regression Model Showing the Association Between the Use of an Anterior Plate (>50% of Cases) and Study Covariates.

Covariate	OR	95% CI	p
<b>AO Spine Region</b>			
Europe and Southern Africa (reference)			
Asia Pacific	2.63	1.49-4.70	<b>0.0009</b>
Latin America	1.46	0.73-2.87	0.2805
North America	25.02	9.31-87.70	<b>&lt;0.0001</b>
Middle East and Northern Africa	2.18	1.09-4.36	<b>0.0272</b>
<b>Specialty</b>			
Orthopaedic surgeon (reference)			
Neurosurgeon	0.34	0.21-0.55	0.8531
Other	0.60	0.09-3.73	0.5753
<b>No use of interbody cages</b>	2.11	0.99-4.71	0.0598

Statistically significant P values are shown in bold.

data. Although providing valuable insights, the reliance on self-reported data within this investigation may introduce response bias. Additionally, some regions, particularly North America, were significantly underrepresented compared to others, such as Europe and Southern Africa and Asia Pacific. The limited data from these regions could skew the results toward the small sample of surgeons who responded to the questionnaire. Altogether, these confounders may impact the reliability of our analyses and affect data interpretation. Therefore, a more comprehensive survey in these underrepresented areas is necessary to better understand their practice patterns and enhance the generalizability of our findings. Furthermore, as a cross-sectional study, this research cannot establish causality or track changes in practice over time.

## Conclusions

In conclusion, the findings of this study highlight the variation in practice patterns and surgeon attitudes regarding use of osteobiologics in ACDF surgery. The data of the survey reveal the need for more formalized and comprehensive training, better awareness of the costs and evidence supporting osteobiologic use, and the dissemination of standardized guidelines. Addressing these issues could lead to improved patient outcomes in ACDF surgeries.

## Acknowledgments

This study was organized and funded by AO Spine through the AO Spine Knowledge Forum Degenerative, a focused group of international spine degeneration experts. AO Spine is a clinical division of the AO Foundation, which is an independent medically-guided not-for-profit organization. Study support was provided directly through the AO Spine Research Department.

## Declaration of Conflicting Interest

The author(s) declared the following potential conflicts of interest with respect to the research, authorship, and/or publication of this article: European Association of Neurosurgical Societies (immediate past-president), Global Neuro Foundation (vice-president); AO Spine Knowledge Forum Degenerative (steering committee and associate members); Depuy Synthes Spine (consulting work); Globus Medical (consulting work, fellowship education support to institution, royalties); Medtronic (consulting fees, royalties, support for attending meetings and/or travel, support to institution); 3M Science (educational consulting), SI Bone (support to institution, royalties), ATEC (support for attending meetings and/or travel, consulting fees), AO Foundation (contracts, support for attending meetings and/or travel), SeaSpine (support to institution), Next Science (support to institution), Motion Metrics (support to institution), NIH SBIR (support to institution), Cerapedics (consulting fees, fellowship education support to institution), AO Spine (consulting fees, support for attending meetings and/or travel, fellowship education support to institution), The Scripps Research Institute (consulting fees), Xenco Medical (consulting fees), North American Spine Society (support for attending meetings and/or travel, committee member, board of directors and executive committee), AO Spine North America (research committee member), Lumbar Spine Research Society (co-chair educational committee), Fehling Instruments GmbH (royalties), Stayble Therapeutics (consulting fees), Mundipharma (consulting fees), Spimplant GmbH (stock options), NC Biomatix (stock options), Stryker (consulting fees), Carlsmed (consulting fees), Alphatec (consulting fees, stock options), Zimvie Spine (royalties), American Orthopaedic Association (committee member), Cervical Spine Research Society (committee member), Scoliosis Research Society (committee member), Empirical Spine (support to institution), International Society for the Study of the Lumbar Spine (support for attending meetings and/or travel, board member), Medsysey (stock options).

## Funding

The author(s) received no financial support for the research, authorship, and/or publication of this article.

## Ethical statement

### Ethical approval

No formal institutional review board approval was required for this study.

### Informed consent

All the participants signed a digital informed consent and agreed on the use of their anonymized responses for research purposes.

## ORCID iDs

- Luca Ambrosio  <https://orcid.org/0000-0003-2424-1274>
- Arun Kumar Viswanadha  <https://orcid.org/0000-0002-6322-4357>
- Zorica Buser  <https://orcid.org/0000-0002-5680-0643>
- Hans Joerg Meisel  <https://orcid.org/0000-0003-3838-1489>
- Hai V. Le  <https://orcid.org/0000-0002-9111-9060>
- Amit Jain  <https://orcid.org/0000-0002-9983-3365>

Andreas K. Demetriades  <https://orcid.org/0000-0002-2004-9448>  
 Sam K. Cho  <https://orcid.org/0000-0001-7511-2486>  
 Patrick C. Hsieh  <https://orcid.org/0000-0002-7206-4842>  
 Tim Yoon  <https://orcid.org/0000-0003-1010-6952>  
 Sathish Muthu  <https://orcid.org/0000-0002-7143-4354>

## Supplemental Material

Supplemental material for this article is available online.

## References

1. Buttermann GR. Anterior cervical discectomy and fusion outcomes over 10 years. *Spine*. 2018;43:207-214. doi:[10.1097/brs.0000000000002273](https://doi.org/10.1097/brs.0000000000002273)
2. Vadalà G, Ambrosio L, De Salvatore S, et al. The role of osteobiologics in augmenting spine fusion in unplated anterior cervical discectomy and fusion compared to plated constructs: a systematic review and meta-analysis. *Glob Spine J*. 2024;14: 43-58. doi:[10.1177/21925682231156865](https://doi.org/10.1177/21925682231156865)
3. Vadalà G, Russo F, Ambrosio L, et al. Biotechnologies and biomaterials in spine surgery. *J Biol Regul Homeost Agents*. 2015;29:137-147.
4. Silber JS, Anderson DG, Daffner SD, et al. Donor site morbidity after anterior iliac crest bone harvest for single-level anterior cervical discectomy and fusion. *Spine*. 2003;28:134-139. doi: [10.1097/0000007632-200301150-00008](https://doi.org/10.1097/0000007632-200301150-00008)
5. Buser Z, Brodke DS, Youssef JA, et al. Synthetic bone graft versus autograft or allograft for spinal fusion: a systematic review. *J Neurosurg Spine*. 2016;25:509-516. doi:[10.3171/2016.1.SPINE151005](https://doi.org/10.3171/2016.1.SPINE151005)
6. Hsieh PC, Chung AS, Brodke D, et al. Autologous stem cells in cervical spine fusion. *Glob Spine J*. 2020;11:950-965. doi:[10.1177/2192568220948479](https://doi.org/10.1177/2192568220948479)
7. Ahn JS, Lee JK, Kim JH. Comparative study of clinical outcomes of anterior cervical discectomy and fusion using autologous bone graft or cage with bone substitute. *Asian Spine Journal*. 2011;5:11. doi:[10.4184/asj.2011.5.3.169](https://doi.org/10.4184/asj.2011.5.3.169)
8. Muthu S, Diniz SE, Viswanathan VK, et al. What is the evidence supporting osteobiologic use in revision anterior cervical discectomy and fusion? *Glob Spine J*. 2024;14:173-178. doi:[10.1177/2192568221133751](https://doi.org/10.1177/2192568221133751)
9. Demetriades AK, Mavrovounis G, Deml MC, et al. What is the evidence surrounding the cost-effectiveness of osteobiologic use in ACDF surgery? A systematic review of the literature. *Glob Spine J*. 2024;14:163S-172S. doi:[10.1177/2192568221148139](https://doi.org/10.1177/2192568221148139)
10. Yoon ST, Konopka JA, Wang JC, et al. ACDF graft selection by surgeons: survey of AOSpine members. *Glob Spine J*. 2017;7: 410-416. doi:[10.1177/2192568217699200](https://doi.org/10.1177/2192568217699200)
11. Fortune business insights. Orthobiologics market size, share & industry analysis, by product type (viscosupplements, bone growth factors, demineralized bone matrix (DBM), synthetic bone substitutes, cellular allograft, allografts, and others), by application (spinal fusion, maxillofacial & dental, soft tissue repair, reconstructive & fracture surgery, and others), by end-user (Hospitals & ASCs, Specialty Clinics, and Others), and Regional Forecast, 2024-2032. 2024;1:1.
12. Meisel HJ, Jain A, Wu Y, et al. AO spine guideline for the use of osteobiologics (AOGO) in anterior cervical discectomy and fusion for spinal degenerative cases. *Glob Spine J*. 2024;14: 6-13. doi:[10.1177/21925682231178204](https://doi.org/10.1177/21925682231178204)
13. Glassman SD, Howard JM, Sweet A, et al. Complications and concerns with osteobiologics for spine fusion in clinical practice. *Spine*. 2010;35:1621-1628. doi:[10.1097/BRS.0b013e3181ce11cc](https://doi.org/10.1097/BRS.0b013e3181ce11cc)
14. James AW, LaChaud G, Shen J, et al. A review of the clinical side effects of bone morphogenetic protein-2. *Tissue Engineering Part B: Reviews*. 2016;22:284-297. doi:[10.1089/ten.teb.2015.0357](https://doi.org/10.1089/ten.teb.2015.0357)
15. Arun-Kumar V, Corluka S, Buser Z, et al. Do osteobiologics augment fusion in anterior cervical discectomy and fusion surgery performed with mechanical interbody devices (polyether ether ketone, carbon fiber, metal cages) and is the fusion rate comparable to that with autograft? A systematic review. *Glob Spine J*. 2024;14:24-33. doi:[10.1177/21925682231188626](https://doi.org/10.1177/21925682231188626)
16. Khan SN, Shahzad H. Osteobiologics and value-based care: challenges and opportunities. *International Journal of Spine Surgery*. 2023;17:S44-S52. doi:[10.14444/8560](https://doi.org/10.14444/8560)
17. Goz V, Rane A, Abtahi AM, et al. Geographic variations in the cost of spine surgery. *Spine*. 2015;40:1380-1389. doi:[10.1097/BRS.0000000000001022](https://doi.org/10.1097/BRS.0000000000001022)
18. Durand WM, Ortiz-Babilonia CD, Badin D, et al. Patient out-of-pocket cost burden with elective orthopaedic surgery. *J Am Acad Orthop Surg*. 2022;1:1. doi:[10.5435/jaaos-d-22-00085](https://doi.org/10.5435/jaaos-d-22-00085)
19. Zale C, Nicholes M, Hu S, et al. Surgical site infection prophylaxis with intra-wound vancomycin powder for uninstrumented spine surgeries: a meta-analysis. *Eur Spine J*. 2023; 32:4259-4264. doi:[10.1007/s00586-023-07897-w](https://doi.org/10.1007/s00586-023-07897-w)
20. Gande A, Rosinski A, Cunningham T, et al. Selection pressures of vancomycin powder use in spine surgery: a meta-analysis. *Spine J*. 2019;19:1076-1084. doi:[10.1016/j.spinee.2019.01.002](https://doi.org/10.1016/j.spinee.2019.01.002)
21. Buser Z, Meisel HJ. Can't see the forest for the trees: a common issue with osteobiologics. *Glob Spine J*. 2023;14:5. doi:[10.1177/21925682231180396](https://doi.org/10.1177/21925682231180396)
22. NASS. Coverage recommendations. <https://www.spine.org/Coverage>. accessed on 21 January 2025.