# 1 Image overlay surgery based on augmented reality:

# 2 a systematic review

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#### 14 Abstract

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Augmented Reality (AR) applied to surgical guidance is gaining relevance in clinical practice. ARbased image overlay surgery (i.e. the accurate overlay of patient-specific virtual images onto the body surface) helps surgeons to transfer image data produced during the planning of the surgery (e.g. the correct resection margins of tissue flaps) to the operating room, thus increasing accuracy and reducing surgery times. We systematically reviewed 76 studies published between 2004 and August 2018 to explore which existing tracking and registration methods and technologies allow healthcare professionals and researchers to develop and implement these systems in-house. Most studies used non-invasive markers to automatically track a patient's position, as well as customised algorithms, tracking libraries or software development kits (SDKs) to compute the registration between patient-specific 3D models and the patient's body surface. Few studies combined the use of holographic headsets, SDKs and user-friendly game engines, and described portable and wearable systems that combine tracking, registration, hands-free navigation and direct visibility of the surgical site. Most accuracy tests included a low number of subjects and/or measurements and did not normally explore how these systems affect surgery times and success rates. We highlight the need for more procedure-specific experiments with a sufficient number of subjects and measurements and including data about surgical outcomes and patients' recovery. Validation of systems combining the use of holographic headsets, SDKs and game engines is especially interesting as this approach allows to easily develop mobile AR applications, thus facilitating the implementation of AR-based image overlay surgery in clinical practice.

- 34 Keywords: Augmented Reality, Mixed Reality, Surgical Guidance, Surgical Navigation, Holographic
- 35 Headsets, Head-Mounted Displays.

#### 1. Introduction

- 37 AR-based image overlay surgery superimposes patient-specific digital data onto the patient's body
- using Augmented Reality (AR), i.e. it augments the real surgical scene by means of computer
- 39 graphics (Azuma, 1997). This approach helps to reduce surgery times, e.g. by preventing the need for
- 40 surgeons to recall image data produced in the planning of the surgery or by facilitating the
- 41 interpretation of 3D data during surgery (Hummelink et al., 2015, Jiang et al., 2018, Khor et al., 2016,
- 42 Kim, Kim & Kim, 2017, Profeta, Schilling & McGurk, 2016, Vávra et al., 2017). It also has the potential
- 43 to reduce intra- and post-operative complications, e.g. by indicating the exact location of high-risk
- 44 anatomical structures adjacent to the surgical site that are not to be injured or facilitating the
- 45 accurate placement of implants (Fritz et al., 2013, Liu et al., 2014). Typically, AR-based image overlay

46 surgery consists of three major steps: 1) tracking, i.e. acquisition of positional information about the 47 patient; 2) registration, i.e. scaling and alignment of the patient-specific imaging data with the 48 previously acquired positional information and; 3) overlay, i.e. projection of the patient-specific 49 digital data onto the patient's body surface using a display device, e.g. a headset. 50 Tracking and registration methods determine key technical aspects of AR-based image overlay 51 surgery systems, e.g. the level of technical skill required to implement and/or use these systems 52 within a surgical setup. A recent review by Eckert et al. (Eckert, Volmerg & Friedrich, 2019) used a 53 large sample of studies obtained from PubMed and Scopus to discuss tracking methods in AR-based 54 medical training and treatment. However, their research does not provide a detailed analysis of the 55 state-of-the-art of AR-based image overlay for surgical guidance. Another recent review by Fida et al. 56 (2018) discussed AR-based image overlay in open surgery. The authors used a single database for 57 their systematic search (PubMed) and excluded studies on neurosurgery, orthopaedics and 58 maxillofacial surgery, which resulted in a fairly small sample of 13 studies. In addition, they did not 59 include a critical reflection of the tracking and registration methods used in their reviewed studies. 60 Our systematic review focuses on AR-based surgical guidance where patient-specific digital data are overlaid onto the patient's body surface (incl. the patient's internal anatomy once exposed during 61 62 open surgery) and in line with the surgeon's view of the surgical site. In contrast to Eckert, Volmerg & Friedrich (2019), our narrower area of study allowed for a detailed analysis and discussion of the 63 64 results across studies that share a particular aim: to guide surgeons by overlaying content on the 65 patient's body surface. For instance, we excluded surgical training, as well as studies on surgical 66 guidance for minimally invasive surgery because this type of surgery presents different tracking and 67 registration challenges than those in open surgery, e.g. tracking markers or anatomical landmarks 68 inside the patient's body using an endoscopic camera (Li et al., 2016). In addition, we included all 69 types of open surgery in our search and used 8 databases, which resulted in a larger sample of 70 studies than in Fida et al. (2018). Finally, we discussed the implications of different registration 71 methods in terms of their application in clinical practice. Other reviews differ from ours in that they 72 cover a particular surgical discipline (Joda et al., 2019, Bertolo et al., 2019, Sayadi et al., 2019, Bosc 73 et al., 2019, Wong et al., 2018) or do not explore the technical aspects of the tracking and 74 registration methods (Contreras López, Navarro & Crispin, 2019, Sayadi et al., 2019, Yoon et al., 75 2018, Kolodzey et al., 2017). 76 The aim of this review is to assess which existing tracking and registration methods and technologies 77 allow healthcare professionals and researchers to develop and implement these systems in-house. 78 As main objectives, we: a) identify the most commonly used tracking methods and the

computational methods that are easiest to implement and; b) explore the registration accuracy of these systems and to what extent they improve surgical outcomes and reduce invasiveness for patients. This work is part of a larger research project which aims to create a methodological and technological framework for AR-based image overlay surgery within the context of reconstructive surgery.

#### 2. Materials and Methods

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This review follows the Preferred Reporting Items for Systematic Review and Meta-Analyses (PRISMA) guidelines (Liberati et al., 2009). The following scientific databases were used for the systematic search in August 2018: Ovid, Medline, Embase, Scopus, Web of Science, PubMed, IEEE (accessed via the University of Aberdeen) and Google Scholar. The search was performed using the following search terms: Augmented Reality AND Image Guided Surgery OR Surgery OR Computer Assisted Surgery AND Tracking OR Registration OR Projection OR Head Mounted Display OR Heads up display OR Smart Glasses OR Autostereoscopic OR Microscopy OR Retinal Displays. Specific and generic terminology as well as alternate spellings and plurals were considered in the search. The full systematic search strategy is provided in the appendix: <u>S1 Table</u>. We considered research on AR-based image overlay surgery published since 2004 when AR was implemented on a mobile device for the first time (Mohring, Lessig & Bimber, 2004). Outcomes were restricted to scientific journal and conference papers written in English and involving animals, humans (including cadaveric material and/or in vivo clinical data belonging to males and females of all ages) and phantom representations. A selection of the retrieved studies was done by one author (LP) through the screening of their titles and abstracts after all authors agreed on the eligibility criteria. The selected studies were classified according to the variables described in Table 1. The experiments conducted by the selected studies were classified according to the Fiducial Registration Error (FRE) and Target Registration Error (TRE) because they were the most common accuracy metrics considered across the reviewed studies. To perform a risk of bias assessment we ranked the individual reviewed studies based on their quality of evidence following the GRADE guidelines (Guyatt et al., 2008): "high" for randomised control trials and "low" for observational studies. Then, an upgrade/downgrade of the resulting level of quality was done based on each study's characteristics: inclusion of accuracy metrics, sample size and inclusion of information about the surgical outcomes. To assess the risk of bias across studies,

we considered the uniformity of the tracking and registration methods and display technologies

used across them. This research did not require the involvement of patients or members of thepublic.

#### Table 1. Variables used to classify the reviewed studies.

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VARIABLE	Description
Surgical task	Surgical step for which the system provided guidance
Surgery type	Surgical procedure for which the system provided guidance
Tracking method	Method used to obtain positional information about the patient
Non-invasive for	The system does not require the use of invasive markers attached to the
patients	patient's body (yes/no).
Registration	Method used to compute the registration between the patient-specific
method	digital data and the patient's body surface
Compact	The system integrates the tracking, registration and display capabilities in a
	single device (yes/no).
Wireless	The system does not require the use of cables within the operating room
	(yes/no).
Surgical site	The system components do not occlude the surgeon's direct view of the
directly visible	surgical site (yes/no).
Hands-free	The surgical team does not need to manipulate the system throughout
tracking	surgery (yes/no).
Stand-alone	The system is presented as a portable program which does not rely on an
application	operating system (yes/no).
Type of display	Type of device used by the system to project the patient-specific digital data
	on the patient's body surface
Includes accuracy	The study includes experiments to measure the registration accuracy of
metrics	their system (yes/no).
N accuracy	Number of accuracy experiments extracted from each reviewed study
experiments	
Fiducial and target	Distance between corresponding real and digital points after registration of
registration errors	the patient-specific digital data with the patient's body. Typically, the FRE is
(FRE and TRE,	measured at points used to set the registration, while the TRE is measured
respectively)	at points other than those used for registration (Fitzpatrick and West, 2001).
Experimental	Subject on which the FRE and TRE were measured
approach	

N subjects	Number of subjects per experiment
N measurements	Number of measurements per experiment
Success rate	The study includes information about the post-operative outcomes
reported	(yes/no).
Surgery time	The study includes information about the time required to perform the
reported	surgery (yes/no).
Long-term study	The study includes monitoring data about the patient's recovery and surgical outcomes (yes/no).
Type of study	Type of study design (randomised control trial or observational study)
Evidence quality	Quality of the evidence provided by the reviewed studies according to GRADE guidelines [21].

#### 3. Results

The systematic search yielded 1352 publications, 724 after removing duplicates (Fig 1). Publications were selected using the following eligibility criteria: 1) the patient-specific digital data were displayed on the patient's body surface (incl. the patient's internal anatomy once exposed during open surgery) either directly (e.g. using conventional projection) or indirectly (e.g. on live images of the patient seen through a tablet) and; 2) the visualisation was in line with the surgeon's view of the surgical site. Therefore, we excluded studies presenting systems which overlaid the patient-specific digital data onto digital scans or images of the patient's internal anatomy (e.g. as in endoscopic procedures), as well as those requiring the surgeon to look away from the surgical site in order to see the digital images (e.g. on a monitor). Among studies on minimally invasive surgery, we included only those in which the tracked features were part of the patient's external anatomy or environment and the patient-specific digital data were overlaid onto the patient's body surface. In total, we selected 76 publications and generated a database (electronic supplementary material: S1 Appendix). These studies covered a variety of surgical tasks (Table 2) and procedures (appendix: S2 Table) showing that some clinical applications had much wider representation within our sample than others.

Fig 1. Flow diagram showing the systematic search strategy used for this review.

Table 2. Classification of reviewed AR-based image overlay surgery studies according to surgical tasks.

Studies	Articles

SURGICAL	%	N	
TASK			
Locate internal	36.84	28	(Maruyama et al., 2018, Zhang, Chen and Liao, 2017, Jiang et al.,
anatomical			2017, Wen, Chng and Chui, 2017, Yang et al., 2018, Sun et al.,
structures,			2017, Scolozzi and Bijlenga, 2017, Drouin S. et al., 2017, Hou et
tumours and			al., 2016, Cabrilo, Schaller and Bijlenga, 2015, Wang et al., 2015,
haematomas			Zhang X., Chen and Liao, 2015, Pauly et al., 2015, Suenaga et al.,
			2015, Yoshino et al., 2015, Kramers et al., 2014, Wang et al.,
			2014, Deng et al., 2014, Wen et al., 2014, Parrini et al., 2014,
			Han et al., 2013, Mahvash and Tabrizi, 2013, Müller M. et al.,
			2013, Kersten-Oertel et al., 2012, Volonte et al., 2011, Tran et
			al., 2011, Sugimoto et al., 2010, Giraldez et al., 2007)
Indicate correct	31.58	24	(Andress et al., 2018, Cutolo et al., 2016, Eftekhar, 2016,
entry points			Fichtinger et al., 2005, Gavaghan et al., 2012, Gibby et al., 2019,
and trajectories			Khan et al., 2006, Krempien et al., 2008, Lee JD. et al., 2010,
of surgical			Liang et al., 2012, Liao et al., 2010, Ma et al., 2018, Ma et al.,
instruments			2017, Martins et al., 2016, Rodriguez et al., 2012, Shamir et al.,
			2011, Si et al., 2018, Suenaga et al., 2013, Vogt, Khamene and
			Sauer, 2006, Wacker et al., 2005, Wang et al., 2016, Wen et al.,
			2013, Wesarg et al., 2004, Wu et al., 2014)
Indicate correct	21.05	16	(Badiali et al., 2014, Besharati Tabrizi and Mahvash, 2015,
soft tissue			Kosterhon et al., 2017, Lin et al., 2016, Lin et al., 2015, Marmulla
resection			et al., 2005, Mischkowski et al., 2006, Mondal et al., 2015,
margins and			Pessaux et al., 2015, Qu et al., 2015, Shao et al., 2014, Sun et al.,
osteotomy			2016, Tang et al., 2017, Wang et al., 2017, Zhu et al., 2016, Zhu
lines			et al., 2011)
Indicate correct	3.95	3	(Ma et al., 2019, Mahmoud et al., 2017, Zeng et al., 2017)
position of			
implants			
Assist more	3.95	3	(He, Liu and Wang, 2016, Hu, Wang and Song, 2013, Wu et al.,
than one			2018)
surgical task			

Indicate	2.63	2	(Huang et al., 2012, Mezzana, Scarinci and Marabottini, 2011)
anatomical			
asymmetry			

#### 3.1. Tracking Methods

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We classified the reviewed studies into the following categories: electromagnetic tracking, optical marker-less tracking and optical marker-based tracking with complex or simple set-up (Fig 2). Most studies used marker-based optical tracking (64%) (Fig 3), e.g. a system which uses a camera to detect the position of a marker fixed to a patient's teeth and, based on this position, projects osteotomy lines onto the patient's skull (Zhu et al., 2016). From these, infrared cameras that detect retroreflective markers were the most commonly used tracking device (41%) (Ma et al., 2019, Maruyama et al., 2018, Si et al., 2018), followed by RGB cameras (20%) to detect 2D images with easily recognisable features (Jiang et al., 2017, Lin et al., 2015, Zhu et al., 2016) or simple shape objects (Cutolo et al., 2016, Sun et al., 2017, Wang et al., 2015). A few studies used marker-less optical tracking (12%) (Gibby et al., 2019, Wu et al., 2018, Zeng et al., 2017), e.g. a camera to detect the contour of the patient's dentition which is matched with its corresponding points on video images of the patient (Wang et al., 2017). Some studies used electromagnetic tracking (3%) (Ma et al., 2018, Martins et al., 2016) or a manual approach (10%) (Eftekhar, 2016, Hou et al., 2016, Pessaux et al., 2015) to detect the patient's position. The remaining studies used alternative methods (Andress et al., 2018, Mahmoud et al., 2017, Scolozzi and Bijlenga, 2017) or did not specify their tracking method (Rodriguez et al., 2012, Sun et al., 2016). A complete list of the reviewed studies classified based on these categories is available in the appendix: S3 Table. Henceforth, the data analysis focuses on the studies using automatic optical tracking (58 studies). Fig 2. Main tracking methods identified in this review: electromagnetic, optical marker-less and optical marker-based with complex or simple set-up. The diagram also shows the devices used for tracking (yellow), registration (green), overlay (orange) or tracking, registration and overlay using a single device (holographic headset). Fig 3. Reviewed studies organised according to their tracking method. Marker-based tracking, use of cameras to detect objects attached to the patient's body; marker-less tracking, superficial body features or a stripy pattern projected onto the patient's body surface; electromagnetic tracking, use of an electromagnetic transmitter to detect sensors placed on a surgical instrument's tip; manual registration, freehand alignment of the patient-specific digital data onto the patient's body surface. EM - electromagnetic; RGB - Red, Green, Blue; RGB-D - Red, Green, Blue and Depth.

#### 3.2. Registration Methods

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Most reviewed studies used custom algorithms to align patient-specific digital data with the patient's position (Ma et al., 2019, Maruyama et al., 2018, Si et al., 2018) (Table 3), e.g. matching two sets of 3D points corresponding to the position of markers on the patient's body and their corresponding points on the patient's scans (Ma et al., 2019). Some studies used computer tracking libraries and/or Software Development Kits (SDKs) (Cutolo et al., 2016, Wang et al., 2016, Zeng et al., 2017), such as OpenCV (Shao et al., 2014), ARToolkit (<a href="http://www.hitl.washington.edu/artoolkit/">http://www.hitl.washington.edu/artoolkit/</a>) (Lin et al., 2016, Qu et al., 2015, Zhu et al., 2016) or Vuforia SDK (https://www.vuforia.com/) (Kramers et al., 2014, Wen, Chng and Chui, 2017). Both ARToolkit and Vuforia SDK provide algorithms to track 2D and 3D feature points on images and define a shared coordinate system between the digital data and the real world (e.g. the patient). They are sometimes used in combination with game engines (e.g. Unity, <a href="https://unity3d.com/">https://unity3d.com/</a> or Unreal, https://www.unrealengine.com/en-US/) and capture devices such as conventional webcams or other RGB/-D camera systems (Jiang et al., 2017, Wu et al., 2018). Game engines with embedded computer tracking libraries and Software Development Kits (SDKs) (e.g. Vuforia SDK) are userfriendly tools that allow to easily develop mobile AR applications which automatically register digital data with real world features. For instance, Wu et al. (2018) used the Vuforia SDK and Unity to deploy the tracking of an image marker placed in the surgical scene. However, their registration strategy also required custom calculations that detect the patient's position. In contrast, Jiang et al. (2017) used ARToolkit and Unity to deploy both the tracking of an image marker and the registration of the patient-specific digital data with the patient's body surface without relying on custom calculations. Only 16% of the reviewed studies used fully integrated platforms (Drouin et al., 2017, Gibby et al., 2019, Sun et al., 2017), e.g. the Brainlab neuronavigation system (Brainlab, Germany).

Table 3. Reviewed studies organised according to the computation method used for automatic optical tracking and registration. Some studies using fully integrated platforms, tracking libraries/SDKs and game engines also developed custom calculation algorithms.

REGISTRATION	Stud	ies	Articles
METHOD	%	N	
Custom	56.90	33	(Badiali et al., 2014, Deng et al., 2014, Giraldez et al., 2007,
algorithms			He, Liu and Wang, 2016, Hu, Wang and Song, 2013, Krempien
			et al., 2008, Lee et al., 2010, Liang et al., 2012, Liao et al.,
			2010, Lin et al., 2015, Ma et al., 2019, Ma et al., 2017,
			Maruyama et al., 2018, Müller et al., 2013, Pauly et al., 2015,

			Shamir et al., 2011, Si et al., 2018, Suenaga et al., 2013,
			Suenaga et al., 2015, Tang et al., 2017, Tran et al., 2011, Vogt,
			Khamene and Sauer, 2006, Wacker et al., 2005, Wang et al.,
			2014, Wang et al., 2015, Wang et al., 2017, Wen et al., 2013,
			Wen et al., 2014, Wu et al., 2014, Yang et al., 2018, Yoshino
			et al., 2015, Zhang, Chen and Liao, 2017, Zhang, Chen and
			Liao, 2015)
Fully integrated	15.52	9	(Cabrilo, Schaller and Bijlenga, 2015, Drouin et al., 2017,
platforms			Gibby et al., 2019, Khan et al., 2006, Kosterhon et al., 2017,
			Mischkowski et al., 2006, Sun et al., 2017, Wesarg et al.,
			2004, Cutolo et al., 2016)
Tracking	20.69	12	(Gavaghan et al., 2012, Huang et al., 2012, Kersten-Oertel et
libraries/SDKs			al., 2012, Kramers et al., 2014, Lin et al., 2016, Qu et al., 2015,
			Shao et al., 2014, Wang et al., 2016, Wen, Chng and Chui,
			2017, Zeng et al., 2017, Zhu et al., 2016, Zhu et al., 2011)
Tracking	3.45	2	(Jiang et al., 2017, Wu et al., 2018)
libraries/SDKs and			
game engines			
Not specified	3.45	2	(Marmulla et al., 2005, Parrini et al., 2014)

# 3.3. Key Aspects of Augmented-Reality-Based Image OverlaySystems

#### **3.3.1.** Ease of use

Most reviewed studies required the set-up of separate pieces of equipment in the operating room (83%), while a minority used compact systems (12%), e.g. those using headsets, smartphones or a microscope with an integrated tracking device (Gibby et al., 2019, Jiang et al., 2017, Sun et al., 2017) (Fig 4). Headsets can be video see-through or optical see-through and display digital data on a screen or on transparent lenses in front of the surgeon's view, respectively. In most cases, the display device occluded the surgeon's view of the surgical site (66%), except for those studies which used optical see-through headsets, smart glasses or projectors (28%) (Gibby et al., 2019, Maruyama et al., 2018, Wu et al., 2018). A minority of studies used hands-free tracking (33%) (Gibby et al., 2019, Ma et al., 2017, Yang et al., 2018), while most required the manipulation of tracking devices (66%). For instance, some systems required the use of a navigation pointer to localise predefined registration

landmarks on the patient's body during surgery (Kosterhon et al., 2017). Only a few studies presented their systems as stand-alone applications (7%), combined with smart glasses (Maruyama et al., 2018), smartphones (Kramers et al., 2014) or holographic headsets (i.e. optical see-through AR headsets that integrate tracking, registration and display capabilities and recognise voice and gesture commands) (Gibby et al., 2019, Wu et al., 2018) (appendix: <u>S4 Table</u>). In addition, most studies relied on hardware with wired connections (84%), while only a few studies used wireless technology such as holographic headsets, smartphones or tablets (Gibby et al., 2019, Sun et al., 2017, Wu et al., 2018). A classification of the reviewed studies according to the display device used is shown in the appendix: <u>S5 Table</u>.

Fig 4. Classification of reviewed automatic optical tracking studies according to system's usability.

#### 3.3.2. Registration Accuracy

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A total of 38 studies on automatic optical tracking (66%) measured the registration accuracy of their system, while the remaining studies did not explore this or measured variables not considered in this review, e.g. the area of tumour successfully removed during AR-based image overlay surgery (Scolozzi and Bijlenga, 2017). In total, we extracted the mean FRE and/or TRE from 44 experiments (Table 4). Most experiments measured the TRE, which has been described as the actual distance between matching real and digital points after registration as it includes all the errors which may occur during the registration process (Fitzpatrick and West, 2001, West et al., 2001). This review shows that many authors achieved TREs between 1-5 mm (52%), e.g. those using computer tracking libraries/SDKs and game engines (Jiang et al., 2017, Wu et al., 2018) and most studies using headsets (Badiali et al., 2014, Cutolo et al., 2016, Gibby et al., 2019, Jiang et al., 2017, Si et al., 2018, Wang et al., 2016, Wu et al., 2018). Some studies achieved a sub-millimetre accuracy (32%), e.g. a study which used a video see-through headset (Lin et al., 2015) and another one using a non-holographic optical see-through headset (Lin et al., 2016). Many reviewed studies included low numbers of subjects and/or measurements in their experiments and only a few were clinical studies (14%), while most measured the registration accuracy on phantoms. Large number of studies did not measure the accuracy of their systems.

Table 4. Classification of experiments according to the registration accuracy and measurement approach. Some articles presented more than one experiment (Maruyama et al., 2018, Wu et al., 2018, Ma et al., 2017, Deng et al., 2014, Giraldez et al., 2007, Wacker et al., 2005). FRE - Fiducial Registration Error; TRE - Target Registration Error; AR - Augmented Reality.

REGISTRATION ACCURACY		Experi	ments	Articles
		%	N	
FRE	<1 mm	11.36	5	(Krempien et al., 2008, Ma et al., 2019, Wang et al., 2014, Wang et al., 2015, Zeng et al., 2017)
	1-5 mm	6.82	3	(Maruyama et al., 2018, Yang et al., 2018, Zhang, Chen and Liao, 2017)
	>5 mm	0	0	-
	Not specified	81.82	36	(Badiali et al., 2014, Cutolo et al., 2016, Deng et al., 2014, Gibby et al., 2019, Giraldez et al.,
				2007, He, Liu and Wang, 2016, Jiang et al., 2017, Khan et al., 2006, Lee et al., 2010, Liang et al.,
				2012, Liao et al., 2010, Lin et al., 2016, Lin et al., 2015, Ma et al., 2017, Maruyama et al., 2018,
				Mischkowski et al., 2006, Qu et al., 2015, Si et al., 2018, Suenaga et al., 2013, Suenaga et al.,
				2015, Wacker et al., 2005, Wang et al., 2016, Wang et al., 2017, Wen et al., 2013, Wen et al.,
				2014, Wen, Chng and Chui, 2017, Wesarg et al., 2004, Wu et al., 2014, Wu et al., 2018, Yoshino
				et al., 2015, Zhu et al., 2016)
TRE	<1 mm	31.82	14	(Giraldez et al., 2007, He, Liu and Wang, 2016, Liao et al., 2010, Lin et al., 2016, Lin et al., 2015,
				Mischkowski et al., 2006, Suenaga et al., 2013, Suenaga et al., 2015, Wang et al., 2014, Wang et
				al., 2015, Wang et al., 2017, Zeng et al., 2017, Zhang, Chen and Liao, 2017)
	1-5 mm	52.27	23	(Badiali et al., 2014, Cutolo et al., 2016, Deng et al., 2014, Gibby et al., 2019, Jiang et al., 2017,
				Krempien et al., 2008, Lee et al., 2010, Liang et al., 2012, Ma et al., 2019, Ma et al., 2017,

				Maruyama et al., 2018, Qu et al., 2015, Si et al., 2018, Wang et al., 2016, Wen et al., 2013, Wen et al., 2014, Wen, Chng and Chui, 2017, Wu et al., 2018, Yoshino et al., 2015, Zhu et al., 2016)
	>5 mm	11.36	5	(Khan et al., 2006, Wacker et al., 2005, Wesarg et al., 2004, Wu et al., 2014)
	Not specified	4.55	2	(Maruyama et al., 2018, Yang et al., 2018)
Experimental approach	Surgery performance	13.64	6	(Deng et al., 2014, Krempien et al., 2008, Maruyama et al., 2018, Mischkowski et al., 2006, Qu et al., 2015, Zhu et al., 2016)
	Surgery simulation on:			
	Phantom	31.82	14	(Cutolo et al., 2016, Gibby et al., 2019, He, Liu and Wang, 2016, Liang et al., 2012, Lin et al., 2016, Lin et al., 2015, Ma et al., 2019, Ma et al., 2017, Si et al., 2018, Wacker et al., 2005, Wen
				et al., 2013, Wen et al., 2014, Wen, Chng and Chui, 2017, Wesarg et al., 2004)
	Animal	6.82	3	(Ma et al., 2017, Wacker et al., 2005, Wu et al., 2014)
	Cadaver	4.55	2	(Khan et al., 2006, Wang et al., 2016)
	Only AR overlay on:			
	Patient	2.27	1	(Suenaga et al., 2015)
	Phantom	38.64	17	(Badiali et al., 2014, Deng et al., 2014, Giraldez et al., 2007, Jiang et al., 2017, Lee et al., 2010,
				Liao et al., 2010, Maruyama et al., 2018, Suenaga et al., 2013, Wang et al., 2014, Wang et al.,
				2015, Wang et al., 2017, Wu et al., 2018, Yang et al., 2018, Yoshino et al., 2015, Zeng et al.,
				2017, Zhang, Chen and Liao, 2017)
	Cadaver	2.27	1	(Giraldez et al., 2007)

N subjects per	< 10	97.73	43	(Badiali et al., 2014, Cutolo et al., 2016, Deng et al., 2014, Gibby et al., 2019, Giraldez et al.,
experiment				2007, He, Liu and Wang, 2016, Jiang et al., 2017, Khan et al., 2006, Krempien et al., 2008, Lee et al., 2010, Liang et al., 2012, Liao et al., 2010, Lin et al., 2016, Lin et al., 2015, Ma et al., 2019, Ma et al., 2017, Maruyama et al., 2018, Mischkowski et al., 2006, Qu et al., 2015, Si et al., 2018, Suenaga et al., 2013, Suenaga et al., 2015, Wang et al., 2016, Wang et al., 2014, Wang et al., 2015, Wang et al., 2017, Wen et al., 2013, Wen et al., 2014, Wen, Chng and Chui, 2017, Wesarg et al., 2004, Wu et al., 2014, Wu et al., 2018, Yang et al., 2018, Yoshino et al., 2015, Zeng et al., 2017, Zhang, Chen and Liao, 2017)
	10-50	2.27	1	(Zhu et al., 2016)
	> 50	0.0	0	-
N measurements per experiment	< 10	50.00	22	(Badiali et al., 2014, Giraldez et al., 2007, He, Liu and Wang, 2016, Jiang et al., 2017, Lee et al., 2010, Liang et al., 2012, Ma et al., 2019, Ma et al., 2017, Mischkowski et al., 2006, Qu et al., 2015, Si et al., 2018, Suenaga et al., 2013, Wang et al., 2014, Wang et al., 2015, Wang et al., 2017, Wu et al., 2018, Yang et al., 2018, Yoshino et al., 2015, Zhang, Chen and Liao, 2017)
	10-50	34.09	15	(Cutolo et al., 2016, Deng et al., 2014, Gibby et al., 2019, Khan et al., 2006, Krempien et al., 2008, Liao et al., 2010, Lin et al., 2015, Maruyama et al., 2018, Wang et al., 2016, Wen et al., 2014, Wen, Chng and Chui, 2017, Wesarg et al., 2004, Wu et al., 2014, Zeng et al., 2017, Zhu et al., 2016)
	> 50	15.91	7	(Deng et al., 2014, Lin et al., 2016, Maruyama et al., 2018, Suenaga et al., 2015, Wacker et al., 2005, Wen et al., 2013)

#### 3.3.3. Surgical Outcomes and Invasiveness for Patients

232 Only few studies compared the surgical success rates (Cutolo et al., 2016, Gibby et al., 2019, Huang 233 et al., 2012, Liao et al., 2010, Lin et al., 2016, Ma et al., 2017, Qu et al., 2015, Si et al., 2018) and 234 times (Khan et al., 2006, Liao et al., 2010, Mischkowski et al., 2006, Müller et al., 2013) with those 235 achieved in conventional surgery. Similarly, only few authors performed long-term studies 236 (Kosterhon et al., 2017). In terms of invasiveness, most marker-based optical tracking studies used 237 non-invasive tracking markers (Giraldez et al., 2007, Huang et al., 2012, Kramers et al., 2014, 238 Krempien et al., 2008, Lee et al., 2010, Maruyama et al., 2018, Wang et al., 2015, Wen et al., 2013, 239 Wen et al., 2014). These markers were attached to the patient (Cutolo et al., 2016, Parrini et al., 240 2014, Si et al., 2018, Sun et al., 2017), a probe that digitises anatomical landmarks (i.e. superficial 241 body features) (Hu, Wang and Song, 2013, Kosterhon et al., 2017, Ma et al., 2017, Tang et al., 2017), 242 a surgical tool (He, Liu and Wang, 2016) or fiducial markers. Fiducial markers are easily identifiable 243 landmarks fixed to the patient's body surface at the time of scanning which allow preserving the 244 spatial relationships between the patient-specific digital data obtained from the scans and the 245 patient's anatomy. Fiducial markers were attached to dental retainers (Ma et al., 2019, Qu et al., 2015, Suenaga et al., 2013, Tran et al., 2011, Yoshino et al., 2015, Zhu et al., 2016, Zhu et al., 2011), 246 247 placed in the surgical scene (Shao P. et al., 2014), or non-invasively attached to the patient 248 (Besharati Tabrizi and Mahvash, 2015, Cutolo et al., 2016, Deng et al., 2014, Drouin et al., 2017, 249 Kersten-Oertel et al., 2012, Liao et al., 2010, Müller et al., 2013, Shamir et al., 2011, Tran et al., 2011, 250 Wu et al., 2014, Yang et al., 2018, Zhang, Chen and Liao, 2017, Zhang, Chen & Liao, 2015, Zhu et al., 251 2016).

#### 3.4. Risk of Bias

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254 Kosterhon et al., 2017, Sun et al., 2017, Zhu et al., 2016, Cabrilo, Schaller and Bijlenga, 2015, Deng et 255 al., 2014, Zhu et al., 2011, Krempien et al., 2008, Giraldez et al., 2007, Mischkowski et al., 2006, 256 Marmulla et al., 2005). Only one reviewed study was a randomised control trial (Qu et al., 2015). 257 Due to their non-inclusion of accuracy metrics, the small sample size in their experiments and/or the 258 lack of information about surgical outcomes, the reviewed case series and reports were downgraded 259 to studies of "very low" quality of evidence, and the randomised control trial was downgraded to 260 "moderate" quality of evidence (electronic supplementary material: S1 Appendix). In addition, a wide variety of tracking and registration methods and display technologies was found across the 261

reviewed studies (<u>Table 3</u> and appendix: <u>S3</u> and <u>S5 Tables</u>).

Most reviewed studies were case series and reports (Maruyama et al., 2018, Tang et al., 2017,

#### 4. Discussion

To the authors' knowledge, this is the first review that: a) identifies the most commonly used tracking and registration methods and technologies that overlay patient-specific digital data onto the patient's body surface and in line with the surgeon's view of the surgical site; b) evaluate the suitability of these methods for their in-house implementation by healthcare professionals and researchers without relying on advanced engineering and/or programming skills and; c) discusses the key challenges of AR-based image overlay surgery.

Our results show that the tracking method most commonly used among the reviewed studies is

marker-based optical tracking, i.e. the use of markers with an easily recognisable pattern to establish a shared coordinate system between the real environment including the patient and the patient-specific 3D dataset (Fig 3). This is in line with the findings by Eckert et al. (Eckert, Volmerg and Friedrich, 2019) who explored a wider area of study: AR-based medical training and treatment. In addition, the registration between the patient-specific digital data and the patient's body surface is normally achieved by using custom calculation algorithms, while the combination of tracking libraries/SDKs and game engines is very recent (Table 3). This review also demonstrates that these systems, which have normally involved the use of several hardware components and cables, do not normally allow the surgeon's direct view of the surgical site or hands-free tracking, and have rarely been presented as stand-alone applications (Fig 4). As key challenges for current AR-based image overlay surgery, we identified the need to validate these systems through more extensive accuracy metrics and to explore approaches that minimise invasiveness for patients.

#### 4.1. Why is Marker-Based Tracking the Commonest Approach?

The use of markers to register patient-specific digital data with the patient's body surface is very common (Fig 3). There are alternatives to using markers, e.g. marker-less optical tracking where anatomical features with well-defined borders (e.g. contour of the patient's dentition) are detected (Suenaga et al., 2015, Wang et al., 2014, Wang et al., 2017). However, the application of marker-less optical tracking is limited as many surgeries do not necessarily involve the exposure of anatomical features with well-defined borders (e.g. soft tissue flap surgery). Similarly, electromagnetic tracking allows the detection of sensors even when they are not visible, e.g. because they are placed in a surgical instrument's tip inside the patient's body. However, this method may compromise surgical accuracy in operating theatres which include several metallic items as magnetic fields are usually affected by metallic artefacts (Poulin and Amiot, 2002). In the absence of anatomical features with

well-defined borders or in environments with metallic items, marker-based optical tracking is a convenient tracking method. This might explain its high prevalence in our reviewed studies.

Two aspects must be considered to prevent an increased risk of intra- and post-operative complications when exploring the use of marker-based tracking: 1) to avoid occlusion of the surgeon's view of the surgical site caused by the markers and; 2) to implement solutions which ensure both an optimal accuracy and low invasiveness for patients. This review shows that there is a variety of options that currently allow the efficient use of non-invasive markers attached to the patients' body surface that minimise their discomfort and facilitate their recovery, e.g. 2D images detected by holographic headsets can be attached to dental splints (Qu et al., 2015, Zhu et al., 2016, Zhu et al., 2011). However, the use of other types of non-invasive markers (e.g. skin adhesives) can lead to a registration mismatch, e.g. due to changes in the soft tissue shape during resection (Jiang et al., 2017).

#### 4.2. What Computational Method is Easiest to Implement?

Traditionally, the development of AR-based image overlay systems has required advanced engineering and programming skills. Fully integrated platforms are highly efficient and easy to implement in the operating room, but also expensive and not suitable for in-house adjustment to particular surgical needs (Drouin et al., 2017). The customisation of AR-based image overlay surgery systems often involves the development of tracking and registration algorithms (Badiali et al., 2014, Wen et al., 2013, Yang et al., 2018) and/or the use of computer tracking libraries and/or SDKs (e.g. OpenIGTLink) (Gavaghan et al., 2012, Huang et al., 2012, Kersten-Oertel et al., 2012, Kramers et al., 2014, Wang et al., 2016, Wen, Chng and Chui, 2017, Zeng et al., 2017). For this reason, this type of development is not available for a wide range of healthcare professionals and researchers. Some reviewed studies overcame this issue by combining computer tracking libraries (e.g. ARToolkits) or SDKs (e.g. Vuforia SDK) with game engines that can be used to create simple mobile AR applications (Andress et al., 2018, Jiang et al., 2017, Wu et al., 2018). In addition, game engines are increasingly becoming more popular due to their improved graphics performance. However, the number of studies using these tools is still relatively small (Table 3).

### 4.3. What are the Benefits of Holographic Headsets?

Holographic headsets are compatible with the previously described tracking and registration methods. Game-based applications using tracking libraries and SDKs can be deployed not only on mobile devices such as smart phones, but also on more specialised displays such as holographic headsets (e.g. Microsoft HoloLens®, https://www.microsoft.com/en-us/hololens). In addition, these

tools provide easy access to algorithms that detect markers (e.g. fiducial markers) on images and align patient-specific digital data with them, i.e. they are compatible with automatic optical tracking. Holographic headsets integrate mobile hardware, a Holographic Processing Unit (HPU) and Depth (RGB-D) cameras (i.e. cameras able to capture both colour and depth information), allowing their use as tracking, registration and display device without relying on an external CPU. AR applications can be loaded into their HPU and used as stand-alone applications. Their RGB-D cameras can be easily set up for marker-based optical tracking by using game engines like Unity (Andress et al., 2018, Si et al., 2018, Wu et al., 2018) and computer tracking software like Vuforia SDK. In addition, their RGB-D cameras can be used to detect surface patterns in the environment (e.g. a patient's body surface) and allow aligning patient-specific 3D models with the patient's body in a fixed position regardless of the user's movement around the room (Gibby et al., 2019). The digital data is overlaid on the headset's transparent lenses without occluding the surgeon's view of the surgical site. They recognise voice and gesture commands, eliminating the need to manipulate tracking devices and allowing hands-free interaction with the digital data (Andress et al., 2018, Jiang et al., 2017, Si et al., 2018, Wu et al., 2018). In summary, the combination of holographic headsets, tracking libraries/SDKs and game-engines allows a wide range of healthcare professionals and researchers to develop simple AR-based image overlay systems in-house, without relying on engineering expertise or commercial providers of fully integrated platforms. In addition, while a wide variety of wearable technology including AR headsets shows promising results in several clinical areas (Kolodzey et al., 2017, Tepper et al., 2017, Keller, State and Fuchs, 2008), holographic headsets are better in facilitating the development of readily available, portable, and easy to set up AR-based image overlay surgery systems which do not alter the surgical workflow significantly (Kramers et al., 2014) (Fig 4). However, studies exploring suitable methodological frameworks for the use of holographic headsets and testing their registration accuracy are very scarce to date (appendix: <a>S5 Table</a>). Part of the reason for this is their fairly recent release (e.g. Microsoft HoloLens® in 2016) and relatively high prices: e.g. Microsoft HoloLens® and Magic Leap® currently cost over \$2000 (developer editions). For this reason and in spite of their advantages, assessing the potential of holographic headsets for their implementation in clinical

#### 4.4. Study Limitations

practice remains a challenge.

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Outcomes from this systematic review show that the number of studies measuring the accuracy of AR-based image overlay surgery systems is low (<u>Table 4</u>), especially if they are analysed separately based on specific characteristics of the system such as its tracking and registration method (<u>Table 3</u>)

and appendix: \$\frac{53 Table}{}\$. Similarly, studies that compare the achieved surgical success rates and times with those of conventional surgery and that include data about the patient's recovery and surgical outcomes in the long-term are scarce in this review. To validate surgical guidance systems that overlay patient-specific digital data onto the patient's body surface (Table 4), it is necessary to perform more clinical studies that include larger samples of subjects and accuracy measurements and that explore the aforementioned variables. For these reasons, most reviewed studies using automatic optical tracking were ranked as "very low" evidence quality (electronic supplementary material: S1 Appendix) and thus we considered that their accuracy estimates remain uncertain. In spite of our restricted eligibility criteria and even though we downsized our sample to automatic optical tracking for the analysis, there was a lack of methodological homogeneity between studies, e.g. due to the wide variety of approaches within each tracking method (appendix: S3 Table), which affects the risk of bias across the reviewed studies. This has also been reported in other reviews with different eligibility criteria, e.g. those reviews focusing on a specific type of surgical procedure (Contreras López, Navarro and Crispin, 2019, Joda et al., 2019) or on wearable technology (Kolodzey et al., 2017). This lack of homogeneity and the low number of studies using common methodological and technological frameworks (Table 4) impeded statistical comparisons between the categories defined in our classifications. Such a statistical analysis would have allowed us to explore potential correlations between registration accuracy and tracking and registration methods and thus make more specific recommendations for improving registration accuracy in future studies. This contrasts with some AR-based guidance tools for minimally invasive surgery such as those for laparoscopy where Eckert et al. (Eckert, Volmerg and Friedrich, 2019) found a high level of research maturity, i.e. they were considered as successfully validated. Incomplete retrieval of relevant publications must also be considered as our search was limited to publications in English. The search, selection and classification of studies was done by the first author only and our qualitative assessments may be biased due to their subjective nature. Finally, research published after August 2018 is not included in our review.

### 5. Conclusions

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AR-based image overlay surgery is becoming more available to healthcare professionals and researchers by combining holographic headsets, computer tracking libraries and/or SDKs and game engines. However, manufacturers and researchers are facing key challenges for the implementation of these systems in clinical practice, such as the need for validation. Current research on AR-based image overlay surgery struggles to provide a sufficient level of registration accuracy for their use in

clinical practice. There is also the need for more clinical studies that include larger numbers of subjects and measurements as well as data about patients' recovery and surgical outcomes. In addition, further research must explore to what extent these systems improve surgery times and success rates and minimise invasiveness for patients. This knowledge would allow manufacturers and researchers to optimise these technologies based on the surgical needs and perform statistical comparisons that facilitate the design of highly efficient systems. Finally, finding a balance between the cost of holographic headsets and their suitability for implementation in clinical practice is important as these novel devices show key benefits: they are portable and wearable, integrate tracking and registration and hands-free navigation and offer direct visibility of the surgical site.

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# 8. Appendix

#### **S1 Table. Search strategy in MEDLINE.**

Search	Search term/s	N publications
1	Surgery, Computer-Assisted/ or Tomography, X-Ray Computed/ or	483962
	augmented reality.mp. or Endoscopy/ or Laparoscopy/	
2	image guided surg\$.mp. or Surgery, Computer-Assisted/	15684
3	1 and 2	15367
4	track\$.tw.	100868
5	registration.tw.	74110
6	fiducial\$.tw.	2519
7	projector.tw.	847
8	projection.tw.	41826
9	head mounted display\$.tw.	446
10	head mounted display\$.mp. or Surgery, Computer-Assisted/	15617
11	head\$ up display\$.tw.	100
12	"Head and Neck Neoplasms"/ or Carcinoma, Squamous Cell/ or	156219
	head\$ up display\$.mp.	
13	autostereoscop\$.tw.	56
14	microscop\$.tw.	537608
15	smart glasses.tw.	26
16	retinal display\$.tw.	14
17	4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14 or 15 or 16	909270
18	3 and 17	15251
19	augmented reality.tw.	839
20	18 and 19	263

### S2 Table. Reviewed studies organised according to surgical procedure.

SURGERY TYPE	Studies		Articles
	%	N	
Neurosurgery	26.32	20	(Cabrilo, Schaller and Bijlenga, 2015, Deng et al., 2014, Drouin
			et al., 2017, Eftekhar, 2016, Hou et al., 2016, Huang et al.,
			2012, Kersten-Oertel et al., 2012, Kramers et al., 2014,
			Krempien et al., 2008, Liao et al., 2010, Mahvash and Tabrizi,
			2013, Maruyama et al., 2018, Shamir et al., 2011, Sun et al.,
			2017, Sun et al., 2016, Besharati Tabrizi and Mahvash, 2015,
			Yang et al., 2018, Yoshino et al., 2015, Zeng et al., 2017, Zhang,
			Chen and Liao, 2015)
Dental,	22.37	17	(Badiali et al., 2014, Lee et al., 2010, Lin et al., 2016, Lin et al.,
craniomaxillofaci			2015, Ma et al., 2019, Marmulla et al., 2005, Mezzana, Scarinci
al and oral			and Marabottini, 2011, Mischkowski et al., 2006, Qu et al.,
			2015, Suenaga et al., 2013, Suenaga et al., 2015, Tran et al.,
			2011, Wang et al., 2014, Wang et al., 2015, Wang et al., 2017,
			Zhu et al., 2011, Zhu et al., 2016)
Assist several	21.05	16	(Cutolo et al., 2016, Fichtinger et al., 2005, Gavaghan et al.,
surgical			2012, Giraldez et al., 2007, Han et al., 2013, He, Liu and Wang,
procedures			2016, Hu, Wang and Song, 2013, Khan et al., 2006, Martins et
			al., 2016, Mondal et al., 2015, Shao et al., 2014, Vogt,
			Khamene and Sauer, 2006, Wacker et al., 2005, Wen, Chng and
			Chui, 2017, Zhang, Chen and Liao, 2017, Wu et al., 2018)
Abdominal	13.16	10	(Mahmoud et al., 2017, Müller et al., 2013, Pessaux et al.,
			2015, Si et al., 2018, Sugimoto et al., 2010, Tang et al., 2017,
			Volonte et al., 2011, Wen et al., 2013, Wen et al., 2014,
			Wesarg et al., 2004)
Orthopaedic	11.84	9	(Andress et al., 2018, Gibby et al., 2019, Kosterhon et al., 2017,
			Liang et al., 2012, Ma et al., 2018, Ma et al., 2017, Pauly et al.,
			2015, Wang et al., 2016, Wu et al., 2014)
Eye	2.63	2	(Rodriguez et al., 2012, Scolozzi and Bijlenga, 2017)
Endovascular	1.32	1	(Parrini et al., 2014)
Perforator flap	1.32	1	(Jiang et al., 2017)

# S3 Table. Classification of reviewed automatic optical tracking studies according to tracking method.

TRACKING METHOD	Studies		Articles	
	%	N		
Marker-based using:				
Infrared camera	40.79	31	(Cabrilo, Schaller and Bijlenga, 2015, Deng et al., 2014,	
			Drouin et al., 2017, Gavaghan et al., 2012, Giraldez et	
			al., 2007, He, Liu and Wang, 2016, Hu, Wang and Song,	
			2013, Huang et al., 2012, Kersten-Oertel et al., 2012,	
			Khan et al., 2006, Kosterhon et al., 2017, Lee et al.,	
			2010, Liang et al., 2012, Liao et al., 2010, Lin et al.,	
			2016, Ma et al., 2019, Ma et al., 2017, Maruyama et al.,	
			2018, Shamir et al., 2011, Si et al., 2018, Suenaga et al.,	
			2013, Tang et al., 2017, Tran et al., 2011, Vogt,	
			Khamene and Sauer, 2006, Wacker et al., 2005, Wang	
			et al., 2016, Wesarg et al., 2004, Yang et al., 2018,	
			Yoshino et al., 2015, Zhang, Chen and Liao, 2017,	
			Zhang, Chen and Liao, 2015)	
RGB camera	19.74 15		(Badiali et al., 2014, Cutolo et al., 2016, Jiang et al.,	
			2017, Kramers et al., 2014, Lin et al., 2015,	
			Mischkowski et al., 2006, Müller et al., 2013, Parrini et	
			al., 2014, Qu et al., 2015, Shao et al., 2014, Sun et al.,	
			2017, Wang et al., 2015, Wu et al., 2014, Zhu et al.,	
			2016, Zhu et al., 2011)	
RGB-D camera	1.32	1	(Wen et al., 2014)	
Projector and	2.63	2	(Krempien et al., 2008, Wen et al., 2013)	
RGB camera				
Marker-less using:				
RGB camera	3.95	3	(Suenaga et al., 2015, Wang et al., 2014, Wang et al., 2017)	
RGB-D camera	6.58	5	(Gibby et al., 2019, Marmulla et al., 2005, Pauly et al.,	
			2015, Wen, Chng and Chui, 2017, Wu et al., 2018)	

Projector and RGB camera	1.32	1	(Zeng et al., 2017)
Electromagnetic	2.63	2	(Ma et al., 2018, Martins et al., 2016)
Manual	10.53	8	(Eftekhar, 2016, Hou et al., 2016, Mahvash and Tabrizi, 2013, Mezzana, Scarinci and Marabottini, 2011, Pessaux et al., 2015, Sugimoto et al., 2010, Besharati Tabrizi and Mahvash, 2015, Volonte et al., 2011)
Other	10.53	8	(Andress et al., 2018, Fichtinger et al., 2005, Han et al., 2013, Mahmoud et al., 2017, Mondal et al., 2015, Rodriguez et al., 2012, Scolozzi and Bijlenga, 2017, Sun et al., 2016)

### 778 S4 Table. Reviewed studies organised according to the system's usability.

USABILITY Studies		es	Articles			
	%	N				
Compact	12.07	7	(Cutolo et al., 2016, Gibby et al., 2019, Giraldez et al., 2007, Jiang			
			et al., 2017, Kramers et al., 2014, Parrini et al., 2014, Sun et al., 2017)			
Wireless	8.62	5	(Gibby et al., 2019, Kramers et al., 2014, Müller et al., 2013, Sun et al., 2017, Wu et al., 2018)			
Surgical site	27.59	16	(Gavaghan et al., 2012, Gibby et al., 2019, Jiang et al., 2017,			
directly visible			Krempien et al., 2008, Liang et al., 2012, Lin et al., 2016, Marmulla			
			et al., 2005, Maruyama et al., 2018, Shao et al., 2014, Si et al.,			
			2018, Wang et al., 2016, Wen et al., 2013, Wen et al., 2014, Wu et			
			al., 2014, Wu et al., 2018, Zeng et al., 2017)			
Hands-free	32.76	19	(Badiali et al., 2014, Cabrilo, Schaller and Bijlenga, 2015, Cutolo et			
tracking			al., 2016, Gibby et al., 2019, Krempien et al., 2008, Lee et al., 2010,			
			Liang et al., 2012, Ma et al., 2017, Marmulla et al., 2005, Pauly et			
			al., 2015, Suenaga et al., 2013, Suenaga et al., 2015, Tran et al.,			
			2011, Wang et al., 2014, Wang et al., 2015, Wang et al., 2017, Wen			
			et al., 2013, Yang et al., 2018, Yoshino et al., 2015)			
Stand-alone	6.90	4	(Gibby et al., 2019, Kramers et al., 2014, Maruyama et al., 2018,			
application			Wu et al., 2018)			

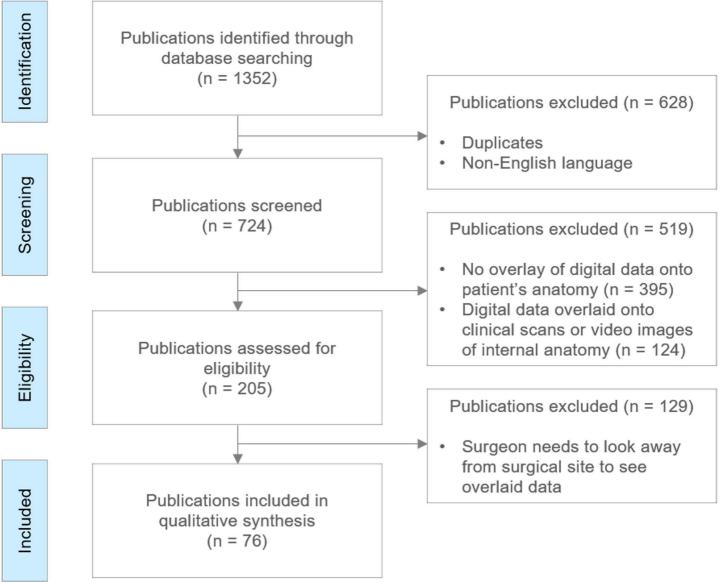
780 S5 Table. Classification of reviewed automatic optical tracking studies according to display device.

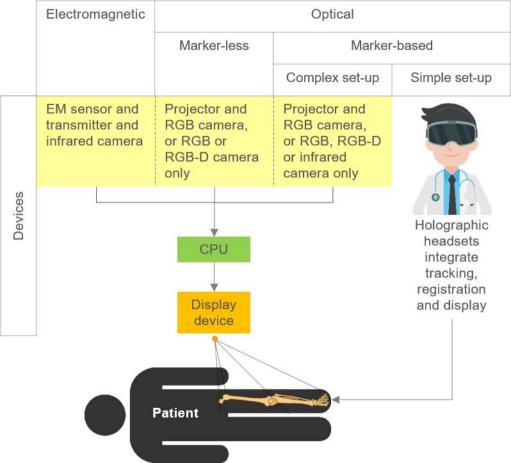
DISPLAY	Studi	es	Articles
	%	N	
Headset			
Video see-through	15.52	9	(Badiali et al., 2014, Cutolo et al., 2016,
			Hu, Wang and Song, 2013, Huang et
			al., 2012, Lin et al., 2015, Parrini et al.,
			2014, Shamir et al., 2011, Vogt,
			Khamene and Sauer, 2006, Wacker et
			al., 2005)
Optical see-through (non-holographic)	5.17	3	(Jiang et al., 2017, Lin et al., 2016,
			Wang et al., 2016)
Optical see-through (holographic)	5.17	3	(Gibby et al., 2019, Si et al., 2018, Wu
			et al., 2018)
Half-silvered mirror	22.41	13	(He, Liu and Wang, 2016, Liao et al.,
			2010, Ma et al., 2019, Ma et al., 2017,
			Pauly et al., 2015, Suenaga et al., 2013,
			Suenaga et al., 2015, Tran et al., 2011,
			Wang et al., 2014, Wang et al., 2015,
			Yang et al., 2018, Zhang, Chen and
			Liao, 2017, Zhang, Chen and Liao,
			2015)
Projector	15.52	9	(Gavaghan et al., 2012, Krempien et al.,
			2008, Lee et al., 2010, Liang et al.,
			2012, Marmulla et al., 2005, Wen et
			al., 2013, Wen et al., 2014, Wu et al.,
			2014, Zeng et al., 2017)
Microscope	8.62	5	(Cabrilo, Schaller and Bijlenga, 2015,
			Drouin et al., 2017, Giraldez et al.,
			2007, Kosterhon et al., 2017, Yoshino
			et al., 2015)
Tablet	8.62	5	(Deng et al., 2014, Mischkowski et al.,
			2006, Müller et al., 2013, Tang et al.,
			2017, Wen, Chng and Chui, 2017)

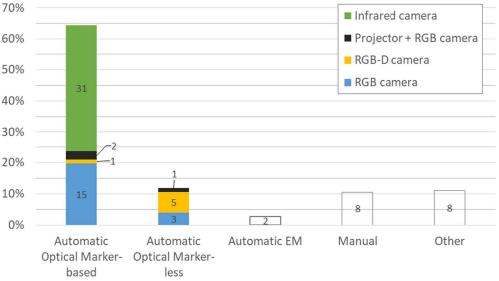
Semi-transparent screen	3.45	2	(Khan et al., 2006, Wesarg et al., 2004)
Smartphone	3.45	2	(Kramers et al., 2014, Sun et al., 2017)
Smart glasses	3.45	2	(Maruyama et al., 2018, Shao et al., 2014)
Video camera screen	1.72	1	(Kersten-Oertel et al., 2012)
Not specified	6.90	4	(Qu et al., 2015, Wang et al., 2017, Zhu et al., 2016, Zhu et al., 2011)

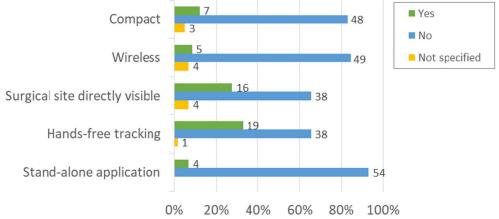
# **9. Supporting information**

- 783 S1 Appendix. Database of reviewed studies categorised according to the variables considered in
- 784 this review.











# **PRISMA 2009 Checklist**

Section/topic	#	Checklist item	Reported on page #
TITLE			
Title	1	Identify the report as a systematic review, meta-analysis, or both.	1
ABSTRACT			
Structured summary	2	Provide a structured summary including, as applicable: background; objectives; data sources; study eligibility criteria, participants, and interventions; study appraisal and synthesis methods; results; limitations; conclusions and implications of key findings; systematic review registration number.	2
INTRODUCTION			
Rationale	3	Describe the rationale for the review in the context of what is already known.	2-4
Objectives	4	Provide an explicit statement of questions being addressed with reference to participants, interventions, comparisons, outcomes, and study design (PICOS).	2-4
METHODS			
Protocol and registration	5	Indicate if a review protocol exists, if and where it can be accessed (e.g., Web address), and, if available, provide registration information including registration number.	4-6
Eligibility criteria	6	Specify study characteristics (e.g., PICOS, length of follow-up) and report characteristics (e.g., years considered, language, publication status) used as criteria for eligibility, giving rationale.	4-6
Information sources	7	Describe all information sources (e.g., databases with dates of coverage, contact with study authors to identify additional studies) in the search and date last searched.	4-6
Search	8	Present full electronic search strategy for at least one database, including any limits used, such that it could be repeated.	4-6
Study selection	9	State the process for selecting studies (i.e., screening, eligibility, included in systematic review, and, if applicable, included in the meta-analysis).	4-6
Data collection process	10	Describe method of data extraction from reports (e.g., piloted forms, independently, in duplicate) and any processes for obtaining and confirming data from investigators.	4-6
Data items	11	List and define all variables for which data were sought (e.g., PICOS, funding sources) and any assumptions and simplifications made.	4-6
Risk of bias in individual studies	12	Describe methods used for assessing risk of bias of individual studies (including specification of whether this was done at the study or outcome level), and how this information is to be used in any data synthesis.	4-6
Summary measures	13	State the principal summary measures (e.g., risk ratio, difference in means).	4-6
Synthesis of results	14	Describe the methods of handling data and combining results of studies, if done, including measures of consistency (e.g., I²) for each meta-analysis.	4-6



## **PRISMA 2009 Checklist**

Section/topic	#	Checklist item	Reported on page #
Risk of bias across studies	15	Specify any assessment of risk of bias that may affect the cumulative evidence (e.g., publication bias, selective reporting within studies).	4-6
Additional analyses	16	Describe methods of additional analyses (e.g., sensitivity or subgroup analyses, meta-regression), if done, indicating which were pre-specified.	-
RESULTS			
Study selection	17	Give numbers of studies screened, assessed for eligibility, and included in the review, with reasons for exclusions at each stage, ideally with a flow diagram.	6
Study characteristics	18	For each study, present characteristics for which data were extracted (e.g., study size, PICOS, follow-up period) and provide the citations.	6-15
Risk of bias within studies	19	Present data on risk of bias of each study and, if available, any outcome level assessment (see item 12).	15
Results of individual studies	20	For all outcomes considered (benefits or harms), present, for each study: (a) simple summary data for each intervention group (b) effect estimates and confidence intervals, ideally with a forest plot.	-
Synthesis of results	21	Present results of each meta-analysis done, including confidence intervals and measures of consistency.	-
Risk of bias across studies	22	Present results of any assessment of risk of bias across studies (see Item 15).	15
Additional analysis	23	Give results of additional analyses, if done (e.g., sensitivity or subgroup analyses, meta-regression [see Item 16]).	-
DISCUSSION	1		
Summary of evidence	24	Summarize the main findings including the strength of evidence for each main outcome; consider their relevance to key groups (e.g., healthcare providers, users, and policy makers).	16-18
Limitations	25	Discuss limitations at study and outcome level (e.g., risk of bias), and at review-level (e.g., incomplete retrieval of identified research, reporting bias).	18-19
Conclusions	26	Provide a general interpretation of the results in the context of other evidence, and implications for future research.	19-20
FUNDING			
Funding	27	Describe sources of funding for the systematic review and other support (e.g., supply of data); role of funders for the systematic review.	20

From: Moher D, Liberati A, Tetzlaff J, Altman DG, The PRISMA Group (2009). Preferred Reporting Items for Systematic Reviews and Meta-Analyses: The PRISMA Statement. PLoS Med 6(6): e1000097. doi:10.1371/journal.pmed1000097

For more information, visit: www.prisma-statement.org.

DOI	Year of publication
10.1007/s11548-018-1814-7	2018
10.1093/ons/opx279	2018
10.3390/s18082505	2018
10.1007/s11517-018-1861-9	2018
10.1007/s11317-018-1801-9 10.1002/rcs.1909	2018
10.1109/ACCESS.2018.2843378	2018
10.1117/1.JMI.5.2.021209	2018
10.1109/SPMB.2017.8257036	2018
10.1007/s11548-017-1634-1	2017
10.1109/TBME.2016.2624632	2017
10.1007/s11548-017-1652-z	2017
10.1097/SAP.000000000001078	2017
10.3390/robotics6020013	2017
10.1097/MD.0000000000008083	2017
10.1016/j.bjoms.2017.08.360	2017
10.1007/s11548-016-1478-0	2017
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10.1007/s11548-016-1444-x	2016
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10.1109/I2MTC.2016.7520404	2016
10.1016/j.wneu.2016.07.047	2016
10.1016/j.wneu.2016.07.107	2016
10.1097/SAP.0000000000000644	2016
10.1007/s00264-015-3028-8	2015
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10.1016/j.compmedimag.2014.11.003	2015
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10.1016/j.compmedimag.2014.06.007	2015
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10.1186/s12880-015-0089-5	2015
10.1016/j.jcms.2014.10.019	2015
10.3171/2014.9.JNS141001	2015
10.2176/nmc.tn.2014-0278	2015
10.1016/j.wneu.2014.12.020	2015
10.1109/TBME.2014.2301191	2014
10.1007/s10439-014-1062-0	2014
10.1159/000354816	2014
10.1016/j.cmpb.2013.12.018	2014
10.1016/j.cmpb.2013.12.021	2014
10.1109/EMBC.2014.6943635	2014
10.1016/j.jcms.2014.09.001.	2014
10.1111/cid.12119	2013
10.3233/978-1-61499-375-9-204	2013
10.1063/1.4830045	2013
10.1007/s00701-013-1668-2	2013
10.1007/s11548-013-0828-4	2013
10.1038/ijos.2013.26	2013
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10.1007/s11548-013-0897-4	2013
10.1109/EMBC.2012.6346203	2012
10.3233/978-1-61499-022-2-225	2012
10.1007/s11548-012-0743-0	2012
10.1088/1748-0221/7/08/P08016	2012
10.1007/s11548-011-0660-7	2011
10.1097/SCS.0b013e31822e8064	2011
10.1007/s00534-011-0385-6	2011
10.1007/978-3-642-23623-5_11	2011
10.1097/PRS.0b013e31820632eb	2011
10.1109/ISBI.2011.5872773	2011
10.1109/TBME.2010.2040278	2010
10.1118/1.3470097	2010
10.1007/s00534-009-0199-y	2009
10.1016/j.ijrobp.2007.10.048	2008
10.1007/s11548-006-0066-0	2007
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10.1007/s11263-006-7938-1	2006
10.1016/j.ijom.2005.05.004	2005
10.1016/j.ics.2005.03.300	2005
10.1109/TBME.2005.851493	2005
10.1117/12.535415	2004

### Title

Head-mounted display augmented reality to guide pedicle screw placement utilizing computed tomography Smart Glasses for Neurosurgical Navigation by Augmented Reality

Visualization

Augmented reality surgical navigation with accurate CBCT-patient registration for dental implant placement distal intramedullary nail interlocking

Mixed Reality Guided Radiofrequency Needle Placement: A Pilot Study

On-the-fly augmented reality for orthopedic surgery using a multimodal fiducial

A Novel Method and System for Stereotactic Surgical Procedures

A surgical robot with augmented reality visualization for stereoelectroencephalography electrode implantation High-quality see-through surgical guidance system using enhanced 3-D autostereoscopic augmented reality study

A Novel Augmented Reality-Based Navigation System in Perforator Flap Transplantation - A Feasibility Study Assisted Surgery

using video-based in situ three-dimensional anatomical modeling: A case report

Removal of recurrent intraorbital tumour using a system of augmented reality

IBIS: an OR ready open-source platform for image-guided neurosurgery

Navigation and Image Injection for Control of Bone Removal and Osteotomy Planes in Spine Surgery

Input System Interface for Image-guided Surgery based on Augmented Reality

On-patient see-through augmented reality based on visual SLAM

Image-guided endoscopic surgery for spontaneous supratentorial intracerebral hematoma

Video see-through augmented reality for oral and maxillofacial surgery

assisted arms - A feasibility study

Procedures in Spine Surgery

App-assisted external ventricular drain insertion

Sensor-fusion based augmented-reality surgical navigation system

iPhone-Assisted Augmented Reality Localization of Basal Ganglia Hypertensive Hematoma

Neuronavigation in Glioma Surgery Involving Eloquent Areas

Effectiveness of a Novel Augmented Reality-Based Navigation System in Treatment of Orbital Hypertelorism system: a pilot study

A high-accuracy surgical augmented reality system using enhanced integral videography image overlay navigation

for tumor resection and sentinel lymph node mapping

Machine learning-based augmented reality for improved surgical scene understanding

Towards cybernetic surgery: robotic and augmented reality-assisted liver segmentectomy

a pilot study

Precise positioning of an intraoral distractor using augmented reality in patients with hemifacial microsomia technique

A Microscopic Optically Tracking Navigation System That Uses High-resolution 3D Computer Graphics

Augmented reality-assisted bypass surgery: embracing minimal invasiveness

D imageoverlay for dental surgery

Designing a wearable navigation system for image-guided cancer resection surgery

Easy-to-use augmented reality neuronavigation using a wireless tablet PC

Hand gesture guided robot-assisted surgery based on a direct augmented reality interface

Real-time advanced spinal surgery via visible patient model and augmented reality system

Augmented reality system for freehand guide of magnetic endovascular devices repositioning

A novel dental implant guided surgery based on integration of surgical template and augmented reality

Evaluation of a mobile augmented reality application for image guidance of neurosurgical interventions

In vivo virtual intraoperative surgical photoacoustic microscopy

A novel augmented reality system of image projection for image-guided neurosurgery

Mobile augmented reality for computer-assisted percutaneous nephrolithotomy

study

A Convenient Method of Video See-through Augmented Reality Based on Image-guided Surgery System

Projection-based visual guidance for robot-aided RF needle insertion

Comparative evaluation of monocular augmented-reality display for surgical microscopes

Augmented reality visualization for guidance in neurovascular surgery

A fluorolaser navigation system to guide linear surgical tool insertion

study on brain-shift estimation

studies

Osteotomy

matter of fashion

Augmented reality system for oral surgery using 3D auto stereoscopic visualization

Augmented reality in oculoplastic surgery: first iPhone application

neurosurgery

3-D augmented reality for MRI-guided surgery using integral videography autostereoscopic image overlay Fast-MICP for frameless image-guided surgery

surgery

brachytherapy

Design and clinical evaluation of an image-guided surgical microscope with an integrated tracking system

Navigation-based needle puncture of a cadaver using a hybrid tracking navigational system

Application of an augmented reality tool for maxillary positioning in orthognathic surgery - A feasibility study evaluation

in the journal International Congress Series

and animals

Image overlay guidance for needle insertion in CT scanner

Accuracy of needle implantation in brachytherapy using a medical AR system - A phantom study

#### Surgical task

Indicate correct entry points and trajectories of surgical instruments Locate internal anatomical structures, tumours and/or haematomas Assist more than one surgical task

Indicate correct position of implants

Indicate correct entry points and trajectories of surgical instruments Indicate correct entry points and trajectories of surgical instruments Indicate correct entry points and trajectories of surgical instruments Locate internal anatomical structures, tumours and/or haematomas Indicate correct position of implants

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Dental, craniomaxillofacial and/or oral Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Neurosurgery Manual  Neurosurgery Automatic optical marker-based [infrared]  Neurosurgery Automatic optical marker-based [infrared]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Assist several surgical procedures Automatic optical marker-based [RGB camera]  Neurosurgery Automatic optical marker-based [RGB-D camera]  Automatic optical marker-based [RGB camera]  Orthopaedic Automatic optical marker-based [RGB camera]  Endovascular Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Neurosurgery Automatic optical marker-based [RGB camera]  Neurosurgery Automatic optical marker-based [RGB camera]  Assist several surgical procedures Other  Neurosurgery Manual  Abdominal Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]		Automatic optical marker-less [RGB-D camera]
Dental, craniomaxillofacial and/or oral Neurosurgery Manual Neurosurgery Automatic optical marker-based [infrared] Neurosurgery Automatic optical marker-based [infrared] Dental, craniomaxillofacial and/or oral Assist several surgical procedures Automatic optical marker-based [RGB camera] Assist several surgical procedures Automatic optical marker-based [RGB-D camera] Neurosurgery Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral	Abdominal	Manual
Dental, craniomaxillofacial and/or oral Neurosurgery Manual Neurosurgery Automatic optical marker-based [infrared] Neurosurgery Automatic optical marker-based [infrared] Dental, craniomaxillofacial and/or oral Assist several surgical procedures Automatic optical marker-based [RGB camera] Assist several surgical procedures Automatic optical marker-based [RGB-D camera] Neurosurgery Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral	Dental, craniomaxillofacial and/or oral	Automatic optical marker-less [RGB camera]
Neurosurgery Automatic optical marker-based [infrared] Neurosurgery Automatic optical marker-based [infrared] Dental, craniomaxillofacial and/or oral Automatic optical marker-less [RGB camera] Assist several surgical procedures Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [Infrared] Abdominal Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [Infrared]	Dental, craniomaxillofacial and/or oral	
Neurosurgery  Dental, craniomaxillofacial and/or oral Assist several surgical procedures Automatic optical marker-based [RGB camera] Assist several surgical procedures Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB-D camera] Abdominal Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [Infrared]	Neurosurgery	Manual
Dental, craniomaxillofacial and/or oral Assist several surgical procedures Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [infrared] Abdominal Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Automatic optical marker-based [Infrared]	Neurosurgery	Automatic optical marker-based [infrared]
Assist several surgical procedures  Neurosurgery  Automatic optical marker-based [RGB camera]  Abdominal  Orthopaedic  Endovascular  Dental, craniomaxillofacial and/or oral  Neurosurgery  Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral  Neurosurgery  Automatic optical marker-based [RGB camera]  Neurosurgery  Automatic optical marker-based [RGB camera]  Assist several surgical procedures  Neurosurgery  Manual  Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral  Automatic optical marker-based [RGB camera]  Automatic optical marker-based [RGB camera]  Automatic optical marker-based [RGB camera]  Automatic optical marker-based [Infrared]	Neurosurgery	Automatic optical marker-based [infrared]
Neurosurgery Automatic optical marker-based [infrared] Abdominal Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Dental, craniomaxillofacial and/or oral	Automatic optical marker-less [RGB camera]
Abdominal Automatic optical marker-based [RGB-D camera] Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Automatic optical marker-based [RGB camera] Automatic optical marker-based [Infrared]	Assist several surgical procedures	Automatic optical marker-based [RGB camera]
Orthopaedic Automatic optical marker-based [RGB camera] Endovascular Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [IRGB camera] Automatic optical marker-based [IRGB camera] Automatic optical marker-based [IRGB camera]	Neurosurgery	Automatic optical marker-based [infrared]
Endovascular Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera]  Neurosurgery Automatic optical marker-based [RGB camera]  Assist several surgical procedures Other  Neurosurgery Manual  Abdominal Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Abdominal	Automatic optical marker-based [RGB-D camera]
Dental, craniomaxillofacial and/or oral Dental, craniomaxillofacial and/or oral Dental, craniomaxillofacial and/or oral Neurosurgery Assist several surgical procedures Neurosurgery Manual Abdominal Dental, craniomaxillofacial and/or oral Automatic optical marker-based [RGB camera] Automatic optical marker-based [RGB camera] Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Orthopaedic	Automatic optical marker-based [RGB camera]
Dental, craniomaxillofacial and/or oral  Neurosurgery  Automatic optical marker-based [RGB camera]  Automatic optical marker-based [RGB camera]  Other  Neurosurgery  Manual  Abdominal  Automatic optical marker-based [RGB camera]  Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral  Automatic optical marker-based [infrared]	Endovascular	Automatic optical marker-based [RGB camera]
Neurosurgery Automatic optical marker-based [RGB camera] Assist several surgical procedures Other Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Dental, craniomaxillofacial and/or oral	Automatic optical marker-based [RGB camera]
Assist several surgical procedures  Neurosurgery  Abdominal  Dental, craniomaxillofacial and/or oral  Other  Manual  Automatic optical marker-based [RGB camera]  Automatic optical marker-based [infrared]	Dental, craniomaxillofacial and/or oral	Automatic optical marker-based [RGB camera]
Neurosurgery Manual Abdominal Automatic optical marker-based [RGB camera] Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Neurosurgery	Automatic optical marker-based [RGB camera]
Abdominal Automatic optical marker-based [RGB camera]  Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Assist several surgical procedures	Other
Dental, craniomaxillofacial and/or oral Automatic optical marker-based [infrared]	Neurosurgery	Manual
	Abdominal	Automatic optical marker-based [RGB camera]
Assist several surgical procedures Automatic optical marker-based [infrared]	Dental, craniomaxillofacial and/or oral	· · · · · · · · · · · · · · · · · · ·
	Assist several surgical procedures	Automatic optical marker-based [infrared]

Abdominal	Automatic optical marker-based [projector and RGB camera]
Eye	Other
Neurosurgery	Automatic optical marker-based [infrared]
Orthopaedic	Automatic optical marker-based [infrared]
Neurosurgery	Automatic optical marker-based [infrared]
Assist several surgical procedures	Automatic optical marker-based [infrared]
Dental, craniomaxillofacial and/or oral	Automatic optical marker-based [RGB camera]
Abdominal	Manual
Dental, craniomaxillofacial and/or oral	Automatic optical marker-based [infrared]
Dental, craniomaxillofacial and/or oral	Manual
Neurosurgery	Automatic optical marker-based [infrared]
Neurosurgery	Automatic optical marker-based [infrared]
Dental, craniomaxillofacial and/or oral	Automatic optical marker-based [infrared]
Abdominal	Manual
Neurosurgery	Automatic optical marker-based [projector and RGB camera]
Assist several surgical procedures	Automatic optical marker-based [infrared]
Assist several surgical procedures	Automatic optical marker-based [infrared]
Dental, craniomaxillofacial and/or oral	Automatic optical marker-based [RGB camera]
Assist several surgical procedures	Automatic optical marker-based [infrared]
Dental, craniomaxillofacial and/or oral	Automatic optical marker-less [RGB-D camera]
Assist several surgical procedures	Automatic optical marker-based [infrared]
Assist several surgical procedures	Other
Abdominal	Automatic optical marker-based [infrared]

It is non-invasive for patients (Y/N)	Registration method	Compact (Y/N)
No	Fully integrated platform	Yes
Yes	Custom calculation algorithms	No
No	Tracking library/SDK and game engine	No
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
No	Tracking library/SDK	No
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No
No	Tracking library/SDK and game engine	Yes
No	Tracking library/SDK	No
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Yes	Fully integrated platform	No
Yes	Fully integrated platform	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
Yes	Fully integrated platform	Yes
No	Custom calculation algorithms	No
No	Tracking library/SDK	No
Yes	Fully integrated platform	Yes
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
Yes	Tracking library/SDK	No
No	Tracking library/SDK  Tracking library/SDK	No
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
No	Custom calculation algorithms	No
Not registered	Not registered	Not registered
No	Custom calculation algorithms	No
Yes	Tracking library/SDK	No
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
No	Fully integrated platform	No
No	Custom calculation algorithms	No
Yes	Tracking library/SDK	No
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No
Yes	•	No
	Custom calculation algorithms	Yes
Yes No	Not specified Custom calculation algorithms	No
No	Custom calculation algorithms	No
	Custom calculation algorithms	Yes
Yes	Tracking library/SDK	
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No

Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Yes	Tracking library/SDK	No
No	Custom calculation algorithms	No
Yes	Tracking library/SDK	No
No	Tracking library/SDK	No
Yes	Tracking library/SDK	No
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	No
Not registered	Not registered	Not registered
Yes	Custom calculation algorithms	No
Yes	Custom calculation algorithms	Yes
No	Fully integrated platform	No
No	Fully integrated platform	No
No	Custom calculation algorithms	No
No	Not specified	No
No	Custom calculation algorithms	No
Not registered	Not registered	Not registered
No	Fully integrated platform	No

Wireless (Y/N)	Surgical site directly visible (Y/N)	Hands-free tracking (Y/N)
Yes	Yes	Yes
No	Yes	No
Yes	Yes	No
No	No	No
Not registered	Not registered	Not registered
No	Yes	No
Not registered	Not registered	Not registered
No	No	Yes
No	Yes	No
No	No	No
No	No	Yes
No	Yes	No
No	No	No
No	No	No
Not registered	Not registered	Not registered
No	No	No
No	No	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
Yes	No	No
No	No	Yes
No	Yes	No
No	No	Yes
Not registered	Not registered	Not registered
No	No	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
No	No	No
No	Yes	No
No	No	No
No	No	Yes
Not registered	Not registered	Not registered
No	No	Yes
Not registered	Not registered	Not registered
No	No	Yes
No	No	No
Not registered	Not registered	Not registered
No	No	Yes
No	No	Yes
No	No	Yes
No	Yes	No
No	No	No
No	Yes	No
No	Yes	No
No	No	No
No	No	Yes
No	No	No
Yes	No	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
Yes	No	No
No	No	Yes
No	No	No

No	Yes	Yes
Not registered	Not registered	Not registered
No	No	No
No	Yes	Yes
No	No	No
No	Yes	No
No	No	No
Not registered	Not registered	Not registered
No	No	Yes
Not registered	Not registered	Not registered
No	No	No
No	No	No
No	No	Yes
Not registered	Not registered	Not registered
No	Yes	Yes
No	No	No
No	Yes	Yes
No	No	No
Not registered	Not registered	Not registered
No	No	No

Stand-alone application (Y/N)	Type of display
Yes	Headset [holographic]
Yes	Smart glasses
Yes	Headset [holographic]
No	Half-silvered mirror
Not registered	Not registered
No	Headset [holographic]
Not registered	Not registered
No	Half-silvered mirror
No	Projector
No	Half-silvered mirror
No	Half-silvered mirror
No	Headset [optical see-through]
No	Tablet
No	Tablet
Not registered	Not registered
No	Microscope
No	Microscope
Not registered	Not registered
Not registered	Not registered
No	Smartphone
No	Not specified
No	Headset [optical see-through]
No	Headset [video see-through]
Not registered	Not registered
No	Half-silvered mirror
Not registered	Not registered
Not registered	Not registered
No	Not registered  Not specified
No	·
	Headset [optical see-through] Half-silvered mirror
No	
No Not remistered	Half-silvered mirror
Not registered	Not registered
No	Half-silvered mirror
Not registered	Not registered
No	Half-silvered mirror
No	Not specified
Not registered	Not registered
No	Microscope
No	Microscope
No	Half-silvered mirror
No	Smart glasses
No	Tablet
No	Projector
No	Projector
No	Headset [video see-through]
No	Headset [video see-through]
No	Headset [video see-through]
Yes	Smartphone
Not registered	Not registered
Not registered	Not registered
No	Tablet
No	Half-silvered mirror
No	Headset [video see-through]
	. 51

No	Projector
Not registered	Not registered
No	Video camera screen
No	Projector
No	Headset [video see-through]
No	Projector
No	Not specified
Not registered	Not registered
No	Half-silvered mirror
Not registered	Not registered
No	Headset [video see-through]
No	Half-silvered mirror
No	Projector
Not registered	Not registered
No	Projector
No	Microscope
No	Semitransparent screen
No	Tablet
No	Headset [video see-through]
No	Projector
No	Headset [video see-through]
Not registered	Not registered
No	Semitransparent screen

Includes accuracy metrics (Y/N)	N experiments	Fiducial registration error (mm)
Yes	1	Not specified
Yes	2	not specified - exp 1, 1-5 - exp 2
Yes	2	Not specified - exp 1, exp 2
Yes	1	<1
Not registered	Not registered	Not registered
Yes	1	Not specified
Not registered	Not registered	Not registered
Yes	1	1-5
Yes	1	<1
Yes	1	1-5
Yes	2	Not specified - exp 1, exp 2
Yes	1	Not specified
Yes	1	Not specified
No	0	Not applicable
Not registered	Not registered	Not registered
No	0	Not applicable
No	0	Not applicable
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
No	0	Not applicable
res	1	Not specified
res	1	Not specified
res	1	Not specified
Not registered	Not registered	Not registered
Yes	1	Not specified
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
Yes	1	Not specified
res Yes	1	Not specified
No	0	Not applicable
√es	1	<1
Not registered	Not registered	Not registered
No	0	Not applicable
Not registered	Not registered	Not registered
∕es	1	Not specified
Yes	1	Not specified
Not registered	Not registered	Not registered
Yes	1	Not specified
No	0	Not applicable
∕es	1	<1
No	0	Not applicable
Yes	2	Not specified - exp 1, exp 2
⁄es	1	Not specified
l'es es	1	Not specified
No	0	Not applicable
l'es	1	Not specified
/es	1	Not specified
No	0	Not applicable
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
No	0	Not applicable
⁄es	1	Not specified
No	0	Not applicable

Yes	1	Not specified
Not registered	Not registered	Not registered
No	0	Not applicable
Yes	1	Not specified
No	0	Not applicable
No	0	Not applicable
No	0	Not applicable
Not registered	Not registered	Not registered
No	0	Not applicable
Not registered	Not registered	Not registered
No	0	Not applicable
Yes	1	Not specified
Yes	1	Not specified
Not registered	Not registered	Not registered
Yes	1	<1
Yes	2	Not specified - exp 1, exp 2
Yes	1	Not specified
Yes	1	Not specified
No	0	Not applicable
No	0	Not applicable
Yes	2	Not specified - exp 1, exp 2
Not registered	Not registered	Not registered
Yes	1	Not specified

Target registration error (mm)	Experimental approach
1-5	Surgery simulation [on phantom]
1-5 - exp 1, not specified - exp 2	Surgery performance - exp 1, AR overlay [on phantom] - exp 2
1-5 - exp 1, exp 2	AR overlay [on phantom] - exp 1, exp 2
1-5	Surgery simulation [on phantom]
Not registered	Not registered
1-5	Surgery simulation [on phantom]
Not registered	Not registered
Not specified	AR overlay [on phantom]
<1	AR overlay [on phantom]
<1	AR overlay [on phantom]
1-5 - exp 1, exp 2	Surgery simulation [on animal - exp 1, on phantom - exp 2]
1-5	AR overlay [on phantom]
1-5	Surgery simulation [on phantom]
Not applicable	Not applicable
Not registered	Not registered
Not applicable	Not applicable
Not applicable	Not applicable
Not registered	Not registered
Not registered	Not registered
Not applicable	Not applicable
<1	AR overlay [on phantom]
<1	Surgery simulation [on phantom]
1-5	Surgery simulation [on phantom]
Not registered	Not registered
<1	Surgery simulation [on phantom]
Not registered	Not registered
Not registered	Not registered
1-5	Surgery performance
1-5	Surgery simulation [on cadaver]
Not applicable	Not applicable
<1	AR overlay [on phantom]
Not registered	Not registered
Not applicable	Not applicable
Not registered	Not registered
<1	AR overlay [on patient]
1-5	Surgery performance
Not registered	Not registered
1-5	AR overlay [on phantom]
Not applicable	Not applicable
<1	AR overlay [on phantom]
Not applicable	Not applicable
1-5 - exp 1, exp 2	Surgery performance - exp 1, AR overlay [on phantom] - exp 2
1-5	Surgery simulation [on phantom]
>5	Surgery simulation [on animal]
Not applicable	Not applicable
1-5	AR overlay [on phantom]
<1	Surgery simulation [on phantom]
	Not applicable
Not applicable	• •
Not registered	Not registered
Not registered	Not registered
Not applicable	Not applicable
<1	AR overlay [on phantom]
Not applicable	Not applicable

1-5	Surgery simulation [on phantom]
Not registered	Not registered
Not applicable	Not applicable
1-5	Surgery simulation [on phantom]
Not applicable	Not applicable
Not applicable	Not applicable
Not applicable	Not applicable
Not registered	Not registered
Not applicable	Not applicable
Not registered	Not registered
Not applicable	Not applicable
<1	AR overlay [on phantom]
1-5	AR overlay [on phantom]
Not registered	Not registered
1-5	Surgery performance
<1 - exp 1, exp 2	AR overlay [on phantom - exp 1, cadaver - exp 2]
>5	Surgery simulation [on cadaver]
<1	Surgery performance
Not applicable	Not applicable
Not applicable	Not applicable
>5 - exp 1, exp 2	Surgery simulation [on animal - exp 1, on phantom - exp 2]
Not registered	Not registered
>5	Surgery simulation [on phantom]

N subjects per experiment	N measurements per experiment
<10	10-50
<10 - exp 1, exp 2	10-50 - exp 1, >50 - exp 2
<10 - exp 1, exp 2	<10 - exp 1, exp 2
<10	<10
Not registered	Not registered
<10	<10
Not registered	Not registered
<10	<10
<10	10-50
<10	<10
<10 - exp 1, exp 2	<10 - exp 1, exp 2
<10	<10
<10	10-50
Not applicable	Not applicable
Not registered	Not registered
Not applicable	Not applicable
Not applicable	Not applicable
Not registered	Not registered
Not registered	Not registered
Not applicable	Not applicable
<10	<10
<10	>50
<10	10-50
Not registered	Not registered
<10	<10
Not registered	Not registered
Not registered	Not registered
10-50	10-50
<10	10-50
Not applicable <10	Not applicable <10
Not registered	Not registered
Not applicable	Not applicable
Not registered	Not registered
<10	>50
<10	<10
Not registered	Not registered
<10	<10
Not applicable	Not applicable
<10	<10
Not applicable	Not applicable
<10 - exp 1, exp 2	10-50 - exp 1, >50 - exp 2
<10	10-50
<10	10-50
Not applicable	Not applicable
<10	<10
<10	10-50
Not applicable	Not applicable
Not registered	Not registered
Not registered	Not registered
Not applicable	Not applicable
<10	<10
Not applicable	Not applicable

<10	>50
Not registered	Not registered
Not applicable	Not applicable
<10	<10
Not applicable	Not applicable
Not applicable	Not applicable
Not applicable	Not applicable
Not registered	Not registered
Not applicable	Not applicable
Not registered	Not registered
Not applicable	Not applicable
<10	10-50
<10	<10
Not registered	Not registered
<10	10-50
<10 - exp 1, exp 2	<10 - exp 1, exp 2
<10	10-50
<10	<10
Not applicable	Not applicable
Not applicable	Not applicable
<10 - exp 1, exp 2	>50 - exp 1, exp 2
Not registered	Not registered
<10	10-50

Success rate reported (Y/N)	Surgery time reported (Y/N)	Long-term study (Y/N)
Yes	No	No
No	No	No
No	No	No
No	No	No
Not registered	Not registered	Not registered
Yes	No	No
Not registered	Not registered	Not registered
No	No	No
No	No	No
No	No	No
Yes	No	No
No	No	No
No	No	No
No	No	No
Not registered	Not registered	Not registered
No	No	No
No	No	Yes
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
-	-	No
No	No No	
No	No	No
Yes	No	No
Yes	No	No
Not registered	Not registered	Not registered
No	No	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
No	No	No
Not registered	Not registered	Not registered
No	No	No
Not registered	Not registered	Not registered
No	No	No
Yes	No	No
Not registered	Not registered	Not registered
No	No	No
Not registered	Not registered	Not registered
Not registered	Not registered	Not registered
No	Yes	No
No	No	No
No	No	No
110	110	110

No	No	No
Not registered	Not registered	Not registered
No	No	No
No	No	No
Yes	No	No
No	No	No
No	No	No
Not registered	Not registered	Not registered
No	No	No
Not registered	Not registered	Not registered
No	No	No
Yes	Yes	No
No	No	No
Not registered	Not registered	Not registered
No	No	No
No	No	No
No	Yes	No
No	Yes	No
No	No	No
No	No	No
No	No	No
Not registered	Not registered	Not registered
No	No	No

Type of study
Not clinical
Clinical [case series]
Not clinical
Not clinical
Not registered
Not clinical
Not registered
Not clinical
Clinical [case report]
Not registered
Not applicable [review - commercial]
Clinical [case report]
Not registered
Not registered
Clinical [case series]
Not clinical
Not clinical
Not clinical
Not registered Not clinical
Not registered
Not registered
Clinical [case series]
Not clinical
Not clinical
Not clinical
Not registered
Not clinical
Not registered
Not clinical
Clinical [randomised control trial]
Not registered
Not clinical
Clinical [case series]
Not clinical
Not clinical
Clinical [case series]
Not clinical
Not registered
Not registered
Not clinical
Not clinical
Not clinical

Not clinical
Not registered
Not clinical
Not clinical
Not clinical
Not clinical
Clinical [case series]
Not registered
Not clinical
Not registered
Not clinical
Not clinical
Not clinical
Not registered
Clinical [case series]
Clinical [case series]
Not clinical
Clinical [case series]
Not clinical
Clinical [case report]
Not clinical
Not registered
Not clinical

Evidence quality
Very low
Very low
Very low
Very low
Not applicable
Very low
Not applicable
Very low
Not applicable
Not applicable
Very low
Not applicable
Not applicable
Very low
Very low
Very low
Very low
Not applicable
Very low
Not applicable
Not applicable
Very low
Very low
Very low
Very low
Not applicable
Very low
Not applicable
Very low
Moderate
Not applicable
Very low
Not applicable
Not applicable
Very low
Very low
Very low

Very low
Not applicable
Very low
Not applicable
Very low
Not applicable
Very low
Very low
Very low
Not applicable
Very low
Not applicable
Very low