Research Article

Educational Use of Animals in Europe Indicates Reluctance to Implement Alternatives

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Abstract

Animals have been considered an indispensable tool to teach about the functioning of living organisms, to obtain skills necessary for practicing human and veterinary medicine, as well as for acquiring skills for caring for and conducting experiments on animals in laboratories. However, the efficacy of this practice has been questioned in recent decades, and societal views have evolved to place a much stronger emphasis on animal welfare and ethics that needs to be reflected in our teaching and training practices. Currently, many alternatives to harmful animal use are available, and it is not clear why thousands of animals continue to be used every year for educational and training purposes. Therefore, this study aimed to identify reasons for the lack of uptake of non-harmful educational and training methods by analyzing recently published non-technical summaries in the EU and EEA Member States and to provide examples of alternatives for specific learning objectives. Results from non-technical summaries from 18 countries spanning the most recent years (2017-2019) revealed that the two main perceived reasons for continued animal use are: 1) the necessity to use a living animal for "proper" learning and 2) the lack of an adequate alternative. We argue that these reasons often do not reflect reality. In conclusion, we consider it is necessary to place a stronger emphasis on engagement with ethical questions that underlie the use of animals and careful consideration of how the learning objectives could be achieved through non-harmful alternatives.

1 Introduction

Animals have been used as anatomy learning tools by artists and scientists for millennia (Hart et al., 2008). Physicians and philosophers in ancient Greece and Rome conducted experiments on live animals to understand animal anatomy, and another upsurge of experimentation on animals occurred during the Renaissance (Bishop and Nolen, 2001). Even today, harmful use of animals remains an integral part of education and training in high schools, universities and the military across the globe (Balls, 2018; Rubeis and Steger, 2018). Animals are killed for use in anatomy dissections and subjected to invasive experiments to demonstrate scientific principles within disciplines such as physiology, biochemistry and pharmacology. They are used also as clinical and surgical skills training models. For instance, pigs are still commonly used in terminal experiments to practice surgical skills in human medicine (Deonarain et al., 2020; Giménez et al., 2020), and rabbits are

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Correspondence: Miriam A. Zemanova (PhD) Animalfree Research Postgasse 15, 3011 Bern, Switzerland (miriam.andela.zemanova@gmail.com) used for training neonatal intubation procedures (Okhovat et al., 2020; Ulmer et al., 2020).

It has been estimated that in any country, 1-10% of animals used in experiments are used for education and training (Akbarsha et al., 2013). Within EU and EEA Member States, thousands of animals are used for educational and training purposes every year (Tab. 1). However, only animals protected by the EU Directive 2010/63 (i.e., vertebrates and cephalopods) are included in the EU and EEA Member States' national statistical reports (EC, 2020). Moreover, not included in the reports are animals that are bred (potentially for other purposes), killed, and their cadavers then used in educational procedures. The actual number of animals used in education and training is therefore likely to be higher. And yet, the use of animals in education has been criticized by both students (Akbarsha, 2007; Oakley, 2013) and educators (Waldau, 2007; Hansen, 2014; Tolbert, 2019). In veterinary education, in particular, any harmful use of animals directly contradicts the commitment of veterinarians to "first, do no harm".

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Tab. 1: Number of animals used for educational and training purposes in the EU and EEA Member States in 2014-2018, according to the national statistical reports

Liechtenstein banned all animal experiments and educational animal use in 1989. Source: European Commission website^a and personal communication with the Norwegian Food Safety Authority.

Country	2014	2015	2016	2017	2018
Austria	3,188	1,925	2,657	3,108	2,889
Belgium	15,178	8,539	8,882	8,051	7,442
Bulgaria	Category not reported				
Croatia	1,188	1,217	1,203	1,428	1,321
Cyprus	Not found				
Czech Republic	7,474	15,673	16,143	7,873	9,172
Denmark	5,745	6,151	5,387	4,463	5,132
Estonia	Category not reported	Category not reported	Category not reported	38	38
Finland	1,738	1,170	1,753	1,606	2,153
France	25,908	28,271	34,280	35,512	41,510
Germany	56,235	55,893	48,941	53,121	55,386
Greece	1,467	1,017	637	1,119	923
Hungary	5,159	1,968	2,688	2,370	2,030
Iceland	Not found				
Ireland	155	389	250	274	343
Italy	1,140	1,042	1,787	1,598	Not yet available
Latvia	150	150	40	22	24
Liechtenstein	Not applicable				
Lithuania	98	44	101	606	1,382
Luxembourg	0	0	0	144	171
Malta	0	0	0	0	Not found
Netherlands	18,240	19,628	14,714	22,074	14,801
Norway	1,065	535	1,097	713	1,027
Poland	4,688	2,688	1,597	1,129	1,117
Portugal	254	3	102	392	228
Romania	2,013	7,498	1,572	1,239	Not yet available
Slovakia	0	8	0	56	30
Slovenia	163	118	53	122	106
Spain	11,149	13,994	12,261	11,785	10,021
Sweden	1,409	3,331	5,155	4,248	4,958
UK	1,814	1,845	1,438	1,235	1,327
TOTAL	148,176	172,044	131,886	162,193	124,345
Countries with missing data	5	5	5	5	7

^a https://ec.europa.eu/environment/chemicals/lab_animals/member_states_stats_reports_en.htm

Moreover, since the implementation of the 3Rs principles (Russell and Burch, 1959) into animal welfare legislation and policy worldwide, the use of animals should be avoided whenever possible (the replacement principle).

Fortunately, there are training methods available that can provide comparable or even better learning outcomes than the use of animals. A broad array of alternatives exists, which has been described in detail elsewhere (Knight, 2007; Zemanova and Knight, 2021). Alternatives include the use of ethically-sourced animal or human cadavers, computer simulations, videos, anatomical models, clinical and surgical skills models, full-body high-fidelity model-driven simulators, virtual reality simulators, non-invasive experimentation on oneself or one's classmates, clinical and surgical rotations, and more. These educational tools usually help overcome educational and training limitations of the more traditional methods (Al-Elg, 2010). They can provide the opportunity to train a wide variety of personnel in technical and non-technical skills without causing harm to animals or real patients. Furthermore, trainees have the possibility to train rare and life-threatening events as many times as necessary (repetitive practice) in a short time. This provides the opportunity to follow controlled training programs, with levels of difficulty adjusted to previous experience, leading to more efficient training (Crofts et al., 2006; Draycott et al., 2006; Guedes et al., 2019). Some simulators and simulation environments provide the possibility to assess performance and debrief after a simulation exercise, based on objective or subjective data. Such features can enhance educational feedback beyond what is generally possible in clinical or scientific practices or when using animals as training subjects (Salas et al., 2009; Grant et al., 2010).

Moreover, the development of non-animal models and animal-free simulation environments for education, training and research is now a stand-alone methodology of similar value to laboratory experiments and clinical trials (Hardman and Ross, 2006; Pawlowski et al., 2018; Labrague et al., 2019). Besides the use of available non-animal methods, new educational and training needs can be met by good curricular design and implementation of potentially new media (Jukes, 2004; Coombs et al., 2017; Pawlowski et al., 2018).

However, the number of animals used for educational and training purposes has remained stable over recent years (Tab. 1), which suggests that such an educational strategy is not yet implemented as standard throughout the EU and EEA Member States and that there may be factors hindering the adoption of non-harmful education and training methods. As a part of the EU Directive 2010/63 on the protection of animals used for scientific purposes, every Member State is required to publish anonymized non-technical summaries (NTSs) of projects using live animals for scientific purposes. A project is defined by the EU Directive 2010/63 as "*a programme of work having a defined scientific objective and involving one or more procedures*", and a procedure means "*any use, invasive or non-invasive, of an animal for experimental or other scientific purposes, with known or unknown*

¹ http://ec.europa.eu/environment/chemicals/lab_animals/nts_en.htm

outcome, or educational purposes, which may cause the animal a level of pain, suffering, distress or lasting harm equivalent to, or higher than, that caused by the introduction of a needle in accordance with good veterinary practice". It is also important to note that experiments that do not affect the physical integrity of the animal can lead to significant distress. For instance, the forced swim test with exhaustion as the end-point – a behavioral experiment used in psychiatric research – has been classified as "severe" (EC, 2009; Strittmatter, 2017).

The requirement for publishing NTSs came into effect for projects authorized after January 1, 2013, except for those authorized under a simplified administrative procedure (Article 42). Each NTS must include information on objectives, predicted harms and benefits, number and species of animals to be used, and a demonstration of adherence to the 3Rs principles (EC, 2013). Therefore, NTSs can provide specific information on what efforts are being made to implement the 3Rs and what obstacles remain for achieving education without harming animals.

The aims of this study were: 1) to elucidate the reasons for continued use of animals for educational and training purposes within the EU and EEA Member States by analyzing recently published NTSs, and 2) to provide an overview of available alternatives for the identified procedures on animals for educational or training purposes.

2 Methods

Search for and analysis of non-technical summaries

First, we searched for the NTSs on the dedicated EU and EEA Member State websites, primarily using the links provided by the EU Commission¹, supplemented with our own search efforts. We only evaluated the NTSs of projects whose sole purpose for using animals was teaching and training. Consequently, the NTSs identified as research projects of PhD or MSc students were not included, as the primary purpose of these projects is basic or applied research, and education is a secondary purpose (Fig. 1). We conducted this survey from October 2019 until May 2020. We decided to assess the most recent years, i.e., 2017-2019, extracting data about the type of procedure on an animal for educational or training purposes, species used, and compliance with the 3Rs principles. In the section on the 3Rs principles, we recorded the reason(s) given to justify the use of animals, i.e., why they could not be replaced. To maintain consistency, the first author assessed all reports.

Review of available educational alternatives

Second, since we suspected that one of the reasons given for continued animal use for teaching and training purposes would be that there are no suitable animal-free models available, we conducted a literature search to identify any alternatives that may already exist for each of the types of procedures on animals for educational or training purposes identified in the

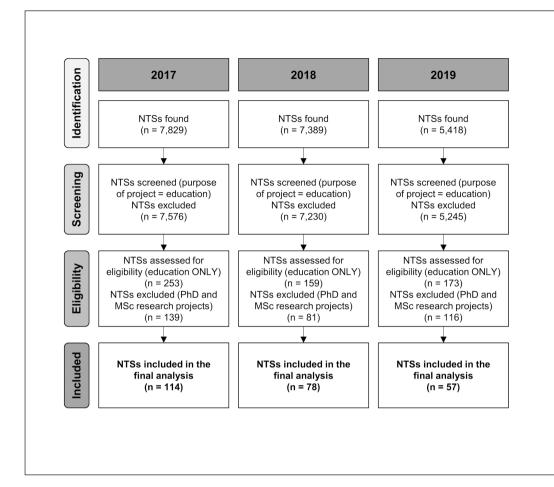


Fig. 1: Steps undertaken during the search for eligible non-technical summaries (NTSs) following the PRISMA guidelines for systematic reviews (Moher et al., 2009) The number of NTSs (n) that were identified, screened, retained or excluded are shown at each stage.

Tab. 2: Key databases that contain information on alternatives in education

Name	lame Description	
InterNICHE Alternatives Database	InterNICHE is the International Network for Humane Education. The network focuses on animal use and alternatives within biological science, medical and veterinary medical education.	http://www.interniche.org/ en/alternatives
NORINA	An English-language database containing information on approximately 3,000 audiovisual aids that may be used as alternatives or supplements to the use of animals in education and training, including dissection alternatives, at all levels from junior school to university. The information in the database has been collected from 1991 onwards.	https://norecopa.no/norina- database
Science Bank	Project of Animalearn (http://www.animalearn.org). Science Bank is a free lending library of humane science products in the US. Links to online resources are also listed.	https://thesciencebank.org/

NTSs. We used search strings containing the type of procedures on animals and the words "alternative", "simulator" or "model" within the Google Scholar bibliographic database. This was chosen for its speed of use, accepting that search results would not be comprehensive and not replicable. Additionally, we consulted several databases of alternative methods for education and training (Tab. 2).

3 Results

3.1 Published non-technical summaries

Liechtenstein banned scientific and educational animal use in 1989 (Kuhse and Singer, 1998). No NTSs could be found for Greece, Malta and Iceland. Croatia, Cyprus, Hungary and Sweden only had NTSs available for the period before 2017. Tab. 3: Number of identified and analyzed non-technical summaries (NTSs) of projects in which animals were used for educational or training purposes in the 31 EU and EEA Member States in 2017-2019 Liechtenstein banned all animal experiments and educational animal use in 1989.

Country	2017	2018	2019	TOTAL
Austria	16	20	8	44
Belgium	4	1	NTSs not yet available	5
Bulgaria	0	NTSs not yet available	NTSs not yet available	0
Croatia	NTSs not yet available	NTSs not yet available	NTSs not yet available	0
Cyprus	NTSs not yet available	NTSs not yet available	NTSs not yet available	0
Czech Republic	16	7	23	46
Denmark	NTSs not searchable	NTSs not searchable	NTSs not searchable	0
Estonia	2	2	2	6
Finland	7	5	2	14
France	11	NTSs not yet available	NTSs not yet available	11
Germany	2	2	3	7
Greece	NTSs not found	NTSs not found	NTSs not found	0
Hungary	NTSs not yet available	NTSs not yet available	NTSs not yet available	0
Iceland	NTSs not found	NTSs not found	NTSs not found	0
Ireland	2	2	0	4
Italy	NTSs not yet available	0	NTSs not yet available	0
Latvia	0	0	1	1
Liechtenstein	Not applicable	Not applicable	Not applicable	0
Lithuania	2	0	NTSs not yet available	2
Luxembourg	0	0	1	1
Malta	NTSs not found	NTSs not found	NTSs not found	0
Netherlands	32	9	0	41
Norway	5	9	10	24
Poland	11	17	5	33
Portugal	0	0	1	1
Romania	NTSs not searchable	NTSs not searchable	NTSs not searchable	0
Slovakia	2	1	1	4
Slovenia	0	1	0	1
Spain	NTSs without dates	NTSs without dates	NTSs without dates	0
Sweden	NTSs not yet available	NTSs not yet available	NTSs not yet available	0
UK	2	2	NTSs not yet available	4
TOTAL	114	78	57	249

The NTSs from Denmark, Romania and Spain were excluded from the analyses due to a format preventing search or due to missing dates. The NTSs from some of the target years have not yet been published by several countries (Tab. 3). In total, for the period 2017-2019, we were able to identify and analyze 249 NTSs purely focused on education and training from 18 out of 31 EU and EEA Member States (Tab. 3; Fig. 1).

3.2 Type of procedures on animals for educational or training purposes, animal

species, and justification for using animals

We identified six general types of procedures on animals associated with educational animal use (Tab. 4; Fig. 2A). Animals were used in 1) behavioral experiments (6 NTSs), 2) experiments in physiology (40 NTSs), 3) insemination and pregnancy diagnosis

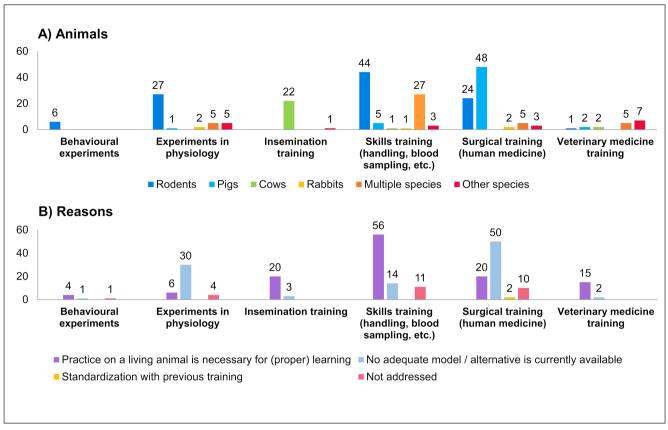


Fig. 2: Results of the analysis of 249 non-technical summaries of animal use for educational and training purposes across 18 EU and EEA Member States in 2017-2019

A) Animal species used for different types of procedures for educational or training purposes (category "Other" includes birds, cats, crustaceans, dogs, foxes, fish, frogs, horses and sheep). B) Reasons reported for why procedures could not be replaced. Please note that the animal procedure type "Skills training" covers primarily laboratory animal courses but might also include NTSs of veterinary medicine training in cases where the exact category was not clear.

training (23 NTSs), 4) skills training (e.g., handling, blood sampling, injection of medications, bronchoscopy training, the practice of euthanasia, 81 NTSs), 5) surgical training in human medicine (82 NTSs) and 6) veterinary medicine training (17 NTSs). The most commonly used species across the types of procedures on animals for educational or training purposes were rodents (102 NTSs), except for insemination training and surgical skills, where cows (23 NTSs) and pigs (48 NTSs), respectively, were used most often.

Among the analyzed NTSs, three main reasons were given for animal use (and not replacing the animal with an alternative): 1) practice on a living animal was deemed necessary for "proper" learning (e.g., because the handling of animals is part of the learning process; 121 NTSs), 2) there was a requirement to use the same approach as in previous trainings to ensure standardization with past teaching (2 NTSs), and 3) no model or alternative currently available was considered adequate (e.g., due to complexity of emergency surgeries; 100 NTSs). Rather surprisingly, 26 NTSs did not include any justification for why replacement was not possible (Fig. 2B). In the following subsections, we list several examples of the analyzed NTSs (Tab. 4), describing how animals were used as well as corresponding available alternatives.

3.3 Examples of available alternatives for the identified types of procedures on animals for educational or training purposes

Our literature search revealed the existence of multiple alternatives for the types of procedures on animals for educational or training purposes identified in the NTSs. Here, we provide only a limited number of examples (Tab. 5) using the same categories used in Table 4 and Figure 2.

3.3.1 Behavioral experiments

Animal behavior can be demonstrated through video documentation (Tab. 5, lines 1-2) of previously conducted experiments (see the examples in Tab. 4, lines 1-2). Other alternatives to using animals in behavioral experiments, in general, are non-harmful interaction with domesticated species, observational studies of sanctuary animals, and field (observational) studies of free-liv-

Line no.	Species	Description of the procedure in the NTS		
Behaviora	l experiments			
1	Rodents	"The aim of the project is to present selected behavioural procedures to students during internships. For this purpose, mild behavioural procedures were selected, presenting an overview of the techniques used to study compounds with antidepressant, anxiolytic and antipsychotic potential. Therefore, a presentation of the most commonly used screening tests such as: SIH, Porsolt test, elevated cross-ma test, tail suspension test, social interactions, and new object recognition tests are planned. The animal will be divided into 2 groups: control and reference group receiving an active dose of a compound with antidepressant (Imipramine) or anxiolytic (Diazepam) or psychotic (MK-801) potential. The overall goal the project is to present students with 'model' behavioural responses so that they can recognize the characteristic behaviour in animals and interpret it correctly."		
2	Rodents	"In addition to the practical work carried out on anesthetized and analgesic animals, students are introduced to behavioural study using various tests carried out in mice. These tests are tools used routinely in the modelling of human pathologies to assess motor, sensory, memory deficits, and test the effectiveness of pharmacological treatments. Students perform five different tests: three motor tests (prehensile traction test, pole test, and footprint test), a memory test (spontaneous alternation), and a resignation test (forced swim test)."		
Physiolog	y experiments			
3	Rodents	"The animal testing labs on rats and mice offered will prepare all students for the acquisition of skills in experimental pharmacology and for understanding the mechanisms of action of drug molecules used in human and veterinary therapy."		
4	Frogs	"The teaching unit allows students to record <i>in vivo</i> the nervous and muscular electrical activities under different experimental conditions. The experiment is done on the sciatic nerve and the gastrocnemius muscle of the frog."		
5	Crustaceans	"Field course for students in zoophysiology, 3 rd -year bachelor. Students will learn how some animals in seawater can live with altered ionic concentration and osmolarity in extracellular fluid when exposed to different salt content. A sample of the extracellular fluid of beach crab in seawater and a sample from a crab after stay in brackish water will be taken."		
Inseminat	ion and pregnancy	diagnosis training		
6	Cows	"Preparatory training for the part of the degree in the insemination of farm animals (cattle)."		
Skills trair	ling			
7	Rodents	"The purpose of this project is to train researchers in a variety of basic procedures that are commonly required within research projects. This is necessary so that the researchers can perform these procedures in a competent manner that minimises distress to the animals."		
8	Rodents	"The aim of this project is to train employees in several biotechnical actions on laboratory animals – mainly standard and some more complex – in the context of building and maintaining animal care and biotechnical skills. Animals will eventually be killed according to legally prescribed methods. The killing car also be part of the training."		
Surgical tr	aining (human me	dicine)		
9	Pigs	"The course aims to train surgeons in minimally invasive surgical techniques. Student surgeons perform various routine operations on anesthetized pigs in order to learn the appropriate skills for interventions on human patients."		
10	Rodents	"This course (Practical Training Course in Microvascular Surgery) has the sole aim and objective of providing suitably skilled surgeons with the highest quality instruction and technical training to allow them to undertake (or support) microvascular surgery on patients in their clinical practice."		
Veterinary	medical training			
11	Rats and rabbits	"Before the actual experiment / manipulation and examination of small mammals (rabbits, rats), the correct examination and other techniques will be presented so that the actual clinical examination and other interventions take place according to the law. For blood collection, intravenous catheter insertion, and urethral catheterization, all animals will be placed under general and analgesics (medetomidine, ketamine, butorphanol) will be administered so that they do not feel pain or fear. Furthermore, endoscopy		

Tab. 4: Examples of procedures on animals for educational or training purposes reported in the non-technical summaries

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Veterinar	Veterinary medical training				
		of the oral cavity, endoscopic examination of the external auditory canal, and vaginoscopy, which are non-invasive or minimally invasive procedures, will be performed. Before laparotomy (access to the abdominal cavity and procedures in the oral cavity), an overdose of anesthetic will be performed and the experimental animal will not regain consciousness. Further surgical procedures will therefore take place on the carcasses."			
12	Cows	"Teaching and assessing clinical skills of undergraduate veterinary students using cattle. The objective is to teach undergraduate veterinary students various techniques including: 1) how to perform epidural anaesthesia (spinal block) and urinary catheterisation, 2) how to administer injections by various routes, 3) how synchronisation of the oestrous cycle (heat period) in cattle can be achieved (this is often performed on commercial farms), 4) how to palpate and ultrasound the female reproductive tract, 5) how to give oral rehydration therapy to adult cows, 6) how to perform a commonly required nerve block, 7) how to perform superovulation (ovulation of a larger than normal number of eggs) and recover embryos (embryos are often transferred from one cow to another which is the main method of moving genetic mating around the world), 8) how to perform various methods of restraint and give oral medication, 9) how to examine the udder and draw milk samples, 10) how to take blood samples At the end, students will be examined on all aspects of clinical competence."			

Tab. 5: Examples of alternatives for each of the types of procedures on animals for educational or training purposes identified in the non-technical summaries

Line no.	Alternative	Description	Source
Behaviora	l experiments	-	
1	Description of the forced swim test Video and protocol		Yankelevitch-Yahav et al., 2015
2	Description of the tail suspension test	Video and protocol	Can et al., 2012
3	Association for the Study of Animal Behaviour: teaching resources	Presentations and videos	https://www.asab.org/higher- education-resources
4	LivingLinks: online learning packs	Presentations, videos and audio recordings	https://living-links.org/resources
5	Methods for Animal Behavior Research	Video	https://www.aza.org/methods- for-animal-behavior-research-dvd
6	Stimulus Response	Video	https://www.stem.org.uk/rx3335
7	Centre for Animal Welfare Science Excellence: video resources	Video	https://www.animalwelfarescience.com/ resources-and-media/video
8	Rabbits and Their Relatives: Health and Management	Computer resource containing text, photos and video	http://www.wildlifeinformation.org
9	Sniffy the Virtual Rat	Laboratory simulation software	http://www.wadsworth.com/ psychology_d/templates/student_ resources/0534633609_sniffy2/sniffy/ main.htm
10	CyberRat	Interactive digital video presentation	http://www.CyberRat.net
Physiolog	y experiments		
11	SimHeart, SimVessel, Drug Laboratory	Physiology and pharmacology experiments in virtual laboratories	http://www.virtual-physiology.com
12	Nerve Physiology	A computer simulation of experiments which may be performed on isolated frog sciatic nerve	http://sheffbp.co.uk/nervephys.html
13	PhysioEx	Laboratory simulation software	https://www.pearson.com.au
14	Monash University	Online interactive simulations	https://ilearn.med.monash.edu.au/ physiology/experiments.html

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	e insemination and pregnancy diagnosis traini		https://www.minitube.com/www.duste/
15	Henryetta	Artificial insemination training cow	https://www.minitube.com/products/ bovine/education-materials/training- model-cow-henryetta-holstein-design
16	Bovine Breeder	Artificial insemination simulator	https://www.realityworks.com/product/ bovine-breeder-artificial-insemination- simulator/
17	Haptic Cow, Haptic Horse	Virtual reality simulator of reproductive tract	http://www.live.ac.uk/haptic-cow
Skills t	raining		
18	Mimicky Mouse®	Mouse simulator for training handling, oral and caudal vein administration	http://www.mitaka-supply.com/mimicky en_index.php
19	Rat training simulator	Model for training handling and procedural competencies	https://www.bioseblab.com/en/ experimental-models/583-rat-training- simulator.html
20	Mimolette Lab Rat Manikin	Simulator for training intubation and blood collection	http://rescuecritters.com/product/ mimolette-mee-moh-let-lab-rat/
21	Critical Care Fluffy / Jerry	Feline / canine manikin for practicing restraint, bandaging, venipuncture, resuscitation	https://rescuecritters.com/product/ critical-care-fluffy/; http://rescuecritters. com/product/critical-care-jerry/
22	Virtual patients	Online learning within veterinary curriculum	Vogt et al., 2019
Surgic	al training – human medicine		
23	SynDaver [®] Surgical Model	Full-body surgical simulator with functional musculoskeletal, cardiovascular, respiratory and gastrointestinal systems	https://syndaver.com/product/ syntissue-surgical-model/
24	TraumaMan	Surgical simulator for training, e.g., chest tube insertion, pneumothorax, tracheostomy, etc.	https://www.simulab.com/products/ traumaman-surgical-simulator
25	LAP Mentor	Laparoscopic simulator	https://simbionix.com/simulators/ lap-mentor/
26	Ligating and Suturing Practice Pad	Pad with a simulated system of vessels to train suturing while a patient is bleeding	https://surgireal.com/collections/ human-simulation/products/ realflow-suture-pad
27	Touch Surgery	Surgical simulation and video platform	https://www.touchsurgery.com/
28	Fundamental Surgery	Virtual reality surgical simulation platform	https://www.fundamentalsurgery.com/
Surgic	al training – veterinary medicine		
29	Canine Dental Surgical Simulator	Simulator for practising canine surgery	https://vetsimulators.com/ caninedentalsimulator/
30	SynDaver	Canine surgical model	https://syndaver.com/product-tag/ 5-veterinary/
31	DASIE	Dog abdominal surrogate for instructional exercise	http://www.dasiesurgery.ca/DASIE/ DASIE.html
32	Microsurgical Developments PVC-Rat	Model to train skills in microsurgery	http://www.microdev.nl/index.php/ products/md-pvc-rat
33	3D Printed Biomimetic Rabbit Airway Simulation Model	Model for nasotracheal intubation training	Oberoi et al., 2020
34	Bovine Vascular Access Model	Model for teaching students a technique for placing catheter in the auricular vein of cattle	Giese et al., 2015

Other (anatomy)				
35	Interspectral	Virtual dissection	https://interspectral.com/what-we-do/ education	
36	Anatomage	Human anatomy 3D visualization tool	https://www.anatomage.com/ anatomage-medical/	
37	zSpace	Virtual reality dissections	https://zspace.com/post-secondary/	
38	eMind	Virtual dissection	https://emindweb.com/products.php	
39	V-Frog	Virtual dissection	http://www.tactustech.com/vfrog/	
40	Interactive Veterinary Education Tool (IVET)	Web-based e-learning platform with 2D and 3D images of body anatomy	Xiberta and Boada, 2019	

ing animals instead of capturing wildlife. Many institutions already provide online learning resources, such as presentations and animal audio and video recordings (Tab. 5, lines 3-10). Examples include the "Stimulus Response" video produced by the Association for the Study of Animal Behaviour, which investigates learning behavior and how animals respond to stimuli in their environments² and videos of farmed animal behavior by The Centre for Animal Welfare Science Excellence³. The computer software resource "Rabbits and Their Relatives: Health and Management" contains information about behavior and behavioral requirements of domestic rabbits (*Oryctolagus cuniculus*) and their wild relatives (rabbits, hares and pikas) through text, photos and video clips⁴.

Additionally, software simulations can provide an interactive training experience. Sniffy the Virtual Rat is a computer program featuring a realistic and interactive laboratory rat in a Skinner box, described as "*Rattus siliconus*". The Sniffy program allows students to design and complete training programs using central behavioral principles such as classical and operant conditioning (Graham et al., 1994). CyberRat is a similar learning tool comprised of video clips of a real laboratory rat in an interactive digital video presentation⁵. Learning principles can also be taught using human demonstrations, which students have reported as being useful and enjoyable (Lewis, 2015).

3.3.2 Experiments in physiology

Drug disposition, pharmacokinetics and mechanisms of action can be easily demonstrated through simulators (e.g., Tab. 5, line 11, and ADAM (Zuna and Holt, 2017); see the example in Tab. 4, line 3). The Sheffield BioScience Programs developed Nerve Physiology, an interactive program that simulates several experiments performed on the frog sciatic nerve (Tab. 5, line 12; see the example in Tab. 4, line 4). Australia's Monash University (Tab. 5, line 14) has established a suite of online interactive practical simulations for physiology classes (Quiroga et al., 2016). There are also other commercially available physiology simulators, e.g., PhysioEx and products from Virtual Physiology (Tab. 5, lines 13 and 11, respectively). If a purpose- or species-specific simulator is not accessible, researchers may use the developed tools to build their own web-based simulators (Šilar et al., 2019) and conduct demonstrations through video documentation of previously conducted experiments (see the example in Tab. 4, line 5).

3.3.3 Insemination and pregnancy diagnosis training

Out of 23 NTSs on insemination and pregnancy diagnosis training, 21 were performed on cows (see the example in Tab. 4, line 6). Breed'n Betsy (Baillie et al., 2005a,b) was one of the first simulators, followed by the more advanced technology implemented in Haptic Cow and Haptic Horse (Baillie et al., 2010). Haptic technology applies anatomically-appropriate pressure to students' fingertips, allowing them to palpate (feel) their way around a virtual simulation (Tab. 5, lines 15-17). The instructor is typically able to observe students' actions on a computer screen and to program the simulation to exhibit different physiological or pathological states (e.g., different stages of pregnancy).

3.3.4 Skills training

Animals are used for skills training of veterinarians, medical doctors, and also personnel working with laboratory animals, mostly rodents. Clinical skills laboratories have been established in many veterinary and medical schools to teach surgical and medical skills, clinical problem-solving, teamwork, and communication, using models, manikins, simulators, and patient actors (Bradley, 2006; Becker, 2007; Grevemeyer and Knight, 2018). Several training simulators are available to train handling skills, venipuncture, intubation or resuscitation techniques that can replace the procedures on animals for educational or training purposes set out in the examples in Tab. 4, lines 7-8. While simulators of rodents seem to be most common, mannequins representing other species are also available (Tab. 5, lines 18-21).

² https://www.stem.org.uk/rx3335

³ https://www.animalwelfarescience.com/resources-and-media/video

⁴ http://www.wildlifeinformation.org

⁵ http://www.cyberrat.net

3.3.5 Human surgical training

Simulation-based training in healthcare has long become an important element between student lectures and proficient clinical practice (Al-Elg, 2010; Meurling et al., 2013; Marker et al., 2019). Historically, medical students trained using cadaveric dissections and clinical practice under the supervision of experienced physicians and surgeons (Wijewickrema et al., 2018). However, today's demand for both student and patient-centered practices and more efficient training programs has led to the development of educational and training programs and environments that allow trainees to acquire competencies adapted to their level and needs (Atesok et al., 2017; Feins et al., 2017; Wijewickrema et al., 2018; Rathod et al., 2019; Villanueva et al., 2019; Aydin et al., 2020). Advanced models using human cadavers that can be connected to artificial blood reservoirs are still used, creating a model that simulates live surgery in terms of bleeding and pulsation (Aboud et al., 2004, 2015; Duhem et al., 2019). Virtual reality simulators (Davies et al., 2013; Cagiltay et al., 2019) and full- and partial-body training simulators such as TraumaMan (Hall et al., 2014), SimMan 3G (Silverplats et al., 2016) or their low-cost alternative SurgeMan (Garcia et al., 2019) provide the technology to fulfil training needs, evaluation of individual technical skills, and team training on non-technical skills. 3D printing has also been implemented to create models for training, for example, cervical laminectomy or pediatric emergency procedures (Kovatch et al., 2020; Tanya and Dubrowski, 2020). These and other simulators are currently used to train surgeons to perform various routine operations in order to learn the appropriate skills for interventions on real patients (Tab. 5, lines 23-25; see the example in Tab. 4, line 9). Tissue suture can be practiced on silicone models (Tab. 5, line 26), some of them with simulated blood (da Silva et al., 2019). A recent systematic review of non-biological microvascular surgery simulators found 20 basic prosthetic models, 20 intermediate models, 13 advanced models, and six virtual reality simulators available for microvascular surgery training (Abi-Rafeh et al., 2019; see example in Tab. 4, line 10).

3.3.6 Veterinary medical training

The use of animals is an integral part of veterinary medical education (see the examples in Tab. 4, lines 11-12). For learning veterinary medical procedures, several e-learning platforms have been developed, e.g., by the University of Girona (Xiberta and Boada, 2019), University of Hannover (Müller et al., 2019), or Freie Universität Berlin (Vogt et al., 2019; Tab. 5, line 22). Multiple surgical simulators are also available (Tab. 5, lines 29-34).

As described in Knight (2011, 2012), humane surgical training courses have been introduced within many veterinary schools. These ideally comprise several stages. First, students learn basic manual skills such as suturing and instrument handling, using

knot-tying boards, plastic organs, and similar models (Tab. 5). Students then progress to simulated surgery on ethically sourced cadavers or full-body or virtual simulators. Major organs may also be perfused with simulated blood via pulsatile pumps (de Souza and Matera, 2015), simulating bleeding and allowing hemostatic practice (control of bleeding). Finally, students observe, assist with, and then perform necessary surgery under close supervision on real patients that benefit from the surgery, similar to the manner in which physicians are trained.

4 Discussion

4.1 Missing information in non-technical summaries

We have identified several concerns with the assessed NTSs in terms of the quality of the information provided. First, there seems to be inconsistency between the numbers reported in the national statistical reports and the numbers reported in the NTSs. For example, Luxembourg reported the use of 144 animals for educational purposes in 2017 (Tab. 1), but we were not able to find any relevant NTSs (Tab. 3). Another example is Germany, which has one of the highest numbers of animals used (2017: 53,121; 2018: 55,386; Tab. 1), but we found only a couple of NTSs describing solely educational procedures (Tab. 3), which were expected to use a total of 24 animals in 2017 and 1,260 in 2018. This could be explained by the fact that in Germany there is no legal obligation for projects involving the use of animals for the primary purpose of training or education to be subjected to the usual authorization procedure. These projects only have to be reported to the authorities⁶. Second, 24 out of 249 NTSs across 7 Member States did not address the required question about the 3Rs and, in particular, how replacement was considered. Lastly, we had difficulties in finding NTSs from some of the Member States. At the time of writing this article, several Member States either did not have a functioning link to their NTSs listed on the European Commission's website or their latest NTSs were more than four years old (Tab. 3). Therefore, there is an issue with irregular and slow publication of NTSs, a problem that has been already reported previously (Taylor et al., 2018). To increase the transparency of animal use, NTSs need to be published regularly, be easily accessible, and strictly follow the template provided by the EU (EC, 2013).

Hopefully, the current lapses in reporting will be remedied by the recently implemented decision EU 2020/569 of the European Commission, "establishing a common format and information content for the submission of the information to be reported by the Member States according to the Directive 2010/63/EU of the European Parliament and the Council on the protection of animals used for scientific purposes and repealing Commission Implementing Decision 2012/707/EU^{°7}.

⁶ https://www.tierschutzbund.de/news-storage/tierversuche/250719-deutschland-laesst-millionen-von-versuchstieren-im-stich/

⁷ https://eur-lex.europa.eu/legal-content/EN/TXT/PDF/?uri=CELEX:32020D0569&from=EN

4.2 Justifications for continued animal use in education and training

The two main reasons given to assert that replacement was not possible, were that "practice on a living animal is necessary for 'proper' learning" and "no adequate model/alternative is currently available" (Fig. 2). Admittedly, it may not be possible to learn certain skills without ever touching a living animal or human. Animal use is certainly needed to practice handling and to gain experience within veterinary education and laboratory animal science, as practice with human beings is necessary for medical education. However, it is important not to inflict unnecessary harm and to promote training programs that could be helpful for animals in addition to the benefits for humans. Veterinary clinical rotations provide one example in which procedures are genuinely intended to benefit animal patients. Another example is provided by animal shelter neutering programs, which are a popular component of humane veterinary surgical courses worldwide as they allow students to gain invaluable experience on some of the most common procedures they will later perform in practice (Richardson et al., 1994; Howe and Slater, 1997). Additionally, the number of unwanted animals killed due to uncontrolled breeding and a subsequent lack of homes is decreased, and neutered animals are more likely to be adopted (Clevenger and Kass, 2003).

We were able to identify several examples of educational and training methods and tools that are currently available for the intended learning outcomes (Tab. 5) and that do not involve the harmful use of animals. There has been a boom in the industry of educational animal-free alternatives: virtual dissection simulations, 3D models, plastinated specimens, videos, or online presentations (Martinsen and Jukes, 2005; Knight, 2007, 2012; Oakley, 2012; Zemanova and Knight, 2021). This diversity of resources allows greater flexibility without compromising the teaching and learning of hands-on skills (Valliyate et al., 2012). Furthermore, with the technologies available today and the rapid evolution of research, veterinary and clinical practices, educational curricula and methods need to be able to promptly adapt to new training needs.

There may be also other reasons for the limited uptake of alternatives that are not mentioned by the authors of the NTSs: resistance from the faculty to change and use alternatives that they themselves did not develop or lack of resources, time or skill for implementation (Dewhurst, 2008; Knight, 2011).

4.3 Are existing models adequate?

There is significant evidence that alternatives can replace many of the identified learning outcomes from procedures on animals for educational or training purposes, such as knowledge and skills acquisition, experimental design, and teamwork (Knight, 2007; Patronek and Rauch, 2007; Dewhurst, 2008; Rosen et al., 2010; Quilici et al., 2015). The first studies on the effectiveness of filmbased instructional activities as dissection alternatives had already emerged by the 1960s (Fowler and Brosius, 1968; De Villiers and Monk, 2005). More recently, Durand et al. (2019) reported that there was no difference in the final grades of students learning about physiology through virtual classes and students learning through animal laboratory classes. Rezende-Filho et al. (2014) re-

ported on a student-centered approach to active learning through the construction of teaching tools in medical physiology. Other studies have found high-fidelity models and simulators realistic, engaging and efficient, and that such training has good transferability to clinical settings (Kinnison et al., 2009; Salas et al., 2009; Grant et al., 2010; Griswold et al., 2012; Marker et al., 2019; Yong et al., 2019; Jaskowski et al., 2020). The systematic review by Mc-Laughlin et al. (2019) revealed that "simulation-based training for trauma resuscitation is associated with improved measures of teamwork, task performance and speed, knowledge and provider satisfaction". Today, more than 99% of all advanced trauma life support training institutions in the USA use simulators exclusively (Pawlowski et al., 2018). In 2006, Dravcott and colleagues showed for the first time that simulation-based training was associated with improved management and clinical outcomes of the trained scenarios (Draycott et al., 2006).

Although there are several alternatives available to learn how to handle rodents and gain procedural competencies (Tab. 5), we noticed that simulation-based training in laboratory animal science is still at an early stage of implementation when compared to healthcare education. For example, research and development in education and training in laboratory animal science is behind in the range of scenarios and the fidelity of simulators available, as well as in the evaluation of learning outcomes, showing that more investment is needed to promote the use of alternatives in this area. However, for any acquisition of skills that will be necessary in future clinical or laboratory practice, simulation-based training without harmful use of animals or humans has proven to be an adaptable methodology to meet any training needs and a suitable practice to improve the knowledge and skills of trainees and the outcomes of patients (Salas et al., 2009; Grant et al., 2010; Griswold et al., 2012; Rubeis and Steger, 2018; Marker et al., 2019; Yong et al., 2019).

4.4 Other concerns associated with continued animal use in education

4.4.1 Harm to the animal

Surgical training in human medicine is normally a non-recovery procedure, i.e., the animal is killed during (normally at the end of) the teaching activity. Whether death is harmful to an animal if inflicted without pain or distress, was often contested historically. Modern conceptualizations of animal welfare, however, understand that good welfare requires not just the avoidance of negative states, but the experience of positive states. Death permanently prevents such positive states and, indeed, the achievement of any other interests animals could seek to fulfil during the remainder of their lives (Balluch, 2006). Accordingly, it is reasonable to conclude that death is one of the most profound harms that can be inflicted (Kaldewaij, 2006; Yeates, 2010; Jensen, 2017), barring exceptional cases such as genuine euthanasia of those faced with severe, ongoing suffering with a poor prognosis for recovery.

Other non-lethal educational practices can also be harmful to an animal. For instance, transrectal palpation training has been shown to increase cortisol levels in cows (Nakao et al., 1994; Cingi et al., 2012; Giese et al., 2018) and horses (Ille et al., 2016; van Vollenhoven et al., 2017), suggesting that the experience is stressful for the animals.

4.4.2 Harm to the student

Veterinary students are drawn to their field of study by their strong interest in animal health and well-being. They can experience stress during their education if they are required to practice painful techniques and terminal surgery (Capaldo, 2004; Gelberg and Gelberg, 2005). Some students may even suffer psychological trauma as a result of engaging in what they perceive as ethically objectionable activities (Capaldo, 2004). The loss of highly qualified people who may forgo a career in science in order not to compromise their values is especially problematic (Capaldo, 2004).

On the other hand, if students are forced to cause suffering to animals during their learning experience, this may lead to "compassion fatigue". This is a condition characterized by a diminished ability to empathize or feel compassion for others (Pereira et al., 2017). Arluke and Hafferty (1996) showed that learning experiences perceived as morally wrong initially lead to ethical uneasiness but ultimately to desensitization through using absolutions justifying the act. Similarly, the decreasing awareness of veterinary students of animal sentience (specifically, the hunger, pain, fear and boredom of dogs, cats and cows) throughout their veterinary courses (Paul and Podberscek, 2000), the decreased likelihood of fourth-year students to provide analgesia when compared to second- or third-year students (Hellyer et al., 1999), and the inhibition of normal development of moral reasoning ability during the four years of veterinary school (Self et al., 1991), have all been revealed within veterinary student cohorts. These are all examples of desensitization-related phenomena. They are actually psychological adaptations to "cognitive dissonance", a discordance between behavior and beliefs (Engel et al., 2020; Pirrone et al., 2019). In this case, the behavior is harmful educational animal use and the belief is that animals are sentient and should not be harmed. Humans normally resolve such dissonance by altering either behavior or beliefs. When altering behavior is not an option - because students fear that refusal to participate could threaten their careers - then beliefs can change, with such desensitization-related phenomena resulting. These adaptations resolve the dissonance and enable previously caring students to withstand what could otherwise be the intolerable psychological stresses that result from being required to harm and kill sentient creatures in the absence of overwhelming necessity (Capaldo, 2004). However, these constitute harms to students and, potentially, risks to their future patients, when veterinarians are subsequently less likely to consider animals as sentient or to warrant appropriate analgesia.

4.4.3 High financial costs

Institutions that use animals for education and training incur costs for caring for animals, such as expenses for veterinary staff, supplies, anesthesia, feeding and disposal. In addition, training procedures may involve the purchase of equipment for each of the student workstations (Graham et al., 1994). Financial barriers related to using live animals were one of the main motivations for creating Sniffy, the Virtual Rat (Graham et al., 1994). Low-technology simulators have the advantage that they can often be easily reproduced by teachers and educators themselves at low cost, which can be particularly important in conditions with a lack of funds and/or when dealing with many students (Adams et al., 2018; Crawford et al., 2019). More expensive tools can also be cost-effective over the longer term. For instance, Gala and Crandall (2019) reported that the introduction of the TraumaMan simulator in advanced trauma life support courses can lead to savings of 25-60 USD per student per year.

4.4.4 Breach of legislation

Article 4 of EU Directive 2010/63 states that: "Member States shall ensure that, wherever possible, a scientifically satisfactory method or testing strategy, not entailing the use of live animals, shall be used instead of a procedure." This Directive has been transposed into national legislation within all EU Member States and was fully implemented by 2013. Our results indicate that in many instances the authors of the NTSs failed to either adequate-ly search for alternatives (e.g., those listed in Tab. 5) or failed to report and implement the results of these searches. This failure to replace animals when there are suitable alternatives available thus indicates non-compliance with the Directive. The "lack of necessity" argument needs to be strictly implemented: "if alternatives exist, animal sacrifice is unnecessary" (De Villiers and Monk, 2005).

5 Conclusion

Between 2014 and 2018, an estimated 124,000 to 172,000 animals were used for educational purposes among the EU and EEA Members States. This is probably an under-estimate, as several countries did not report figures. In view of our results, the use of a large number of these animals could likely have been avoided. Our data appears to indicate that the potential for implementing alternatives has not been fully met and that considerable room for improvement exists. We hope this study will serve as an incentive for educators, universities and funding agencies to support the development and implementation of non-harmful alternatives and curricula in life sciences education and training. We also hope this study will help governments and competent authorities to put in place national policies to replace the use of animals for the purposes of education and training, as well as to effectively evaluate possible replacement alternatives for projects aiming at using animals for educational and training purposes. Wider discussions about and promotion of these alternatives in education and training are crucial for establishing a culture of life and health sciences education that is ethical and compassionate.

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Conflict of interest

The authors declare that they have no conflicts of interest.

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