



# Article Animal Research in Spain: A Study of Public Perception and Attitudes

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**Simple Summary:** Although animals are extensively used in experimentation for diverse purposes, ethical concerns have led to reluctance among a certain sector of the population. To find out the opinions of the Spanish population regarding animal research, we conducted a survey, obtaining responses from more than 800 people. While the majority were in favor, some groups were more prone to be against animal experimentation. In addition, we observed that people's opinions could be altered by reading media reports. We also observed that people did not know about the handling of animals or laws concerning animal testing. Furthermore, we found that people were against the use of animals for non-medical purposes. In general, the survey showed the variety of people's opinions and their concern for animal welfare.

**Abstract:** Since the time of Hippocrates in the 4th century BC, animal research has been extensively used for various purposes up to the present day. However, the use of animals for research has also been controversial for a long time. We report the findings of a public, online questionnaire-based survey designed to assess the opinions of a sample of Spanish society regarding animal research. Demographic data and opinions were obtained from 806 respondents. The results indicated a high level of acceptance of animal research (73.1%). However, certain factors, such as completing the questionnaire immediately after a reading negative media report (OR = 2.41; 95%CI: 1.64–3.54; *p* < 0.001), being a woman (OR = 1.77; 95%CI: 1.24–2.53; *p* = 0.002) or having a non-scientific background (OR = 2.47; 95%CI: 1.76–3.47; *p* < 0.001), were associated with a tendency towards a more negative opinion. The opinions seemed to be influenced by gender, education level and by protest incidents reported in the media. Our results also indicate that a lot of information regarding animal welfare, such as care and handling protocols, along with legislation was unknown to individuals. Further, a growing popularity of companion species and opposition to animal experimentation for non-biomedical purposes were reflected in the responses obtained. The use of animal sfor research purposes emerged as a sensitive social issue in terms of concerns about animal ethics and welfare.

Keywords: animal research; gender; education level; media incident; survey

## 1. Introduction

Since the time of Hippocrates in the 4th century BC, animal research has sought to expand scientific knowledge and resulted in many advances in fields such as biomedicine [1,2]. Effectively, the many species of animals used in research [3,4] have become essential tools to understand and characterize human physiology in terms of both health and disease [5–8].



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**Copyright:** © 2023 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (https:// creativecommons.org/licenses/by/ 4.0/). Also, an estimate of 70% of emerging infectious diseases have animal origins [9]. It is important to emphasize the relevance of animal experimentation in the study of animal diseases, especially zoonotic diseases, and its vital role in Public Health. In addition, the use of new technologies, such as Geographic Information Systems (GIS) and Remote Sensing, can be essential for their management and planning towards the One Health perspective [10,11]. However, some authors claim that animal testing is insufficient to predict clinical outcomes among humans, and they question the reliability of many animal test results. They consider many tests to be a waste of economic resources and harmful to animals [12].

In each country or region, scientific studies involving the use of animals are regulated by different laws: Directive 2010/63/EU of the European Parliament and European Council [13] in Europe, the Animal Welfare Act (AWA) [14] in the United States (protects all warm-blooded animals in research except rats, mice and birds bred), and the Animal Welfare Act [15] in the UK, along with similar regulations in other countries with developed scientific, technological systems. To try to ensure the welfare of the animals used for research purposes, all legislation is based on the principle of the 3Rs defined by Russell and Burch (Replace, Reduce and Refine) [16]. The field of alternative methods to animal experimentation has been growing and diversifying in recent years and has impacted many other disciplines. In this context, the term "alternative methods" is no longer precisely defined and has been replaced by the term, "3R methods" [17]. According to European Directive 2010/63/EU [13], all personnel working with experimental animals must be educated in how to work with animals. In many countries, courses have been established or are still in process to meet this requirement and to be educated and regularly updated on the advances being made with respect to the 3Rs. Due to the complexity of this diverse field, it is mandatory that experts share their knowledge to develop comprehensive teaching programs, gain a clear understanding of the advantages and limitations of the 3Rs approaches and how to convey them to all population target groups [18].

There are numerous limitations regarding animal experimentation such as problems with the translation and extrapolation of results to humans [19,20]. The proper selection of animal species for research is one of the main pillars for any subsequent clinical translation [21,22]. With advancing scientific knowledge and technological progress, alternative models of biomedical research are now emerging, helping to replace experimental animal models [6,23].

Owing to welfare concerns, the use of animals in research has become a controversial issue at political, social, and economic levels. National and international opinion surveys have reflected different points of view in different populations, such as healthcare professionals and veterinarians [24–27]. In a public survey conducted in Japan in 2019 [27], 51–57% of interviewees mentioned they considered animal experiments to be cruel and painful. However, 55–62% stated that animal experimentation is necessary for advances in human medicine and to guarantee the health and safety of people. In another public survey on the use of laboratory animals administered to US medical students and physicians in 2016 [25], there was widespread opposition to using animals in research based in expectations and preferences that alternative models and methods should be used.

According to an opinion survey in 2001 [26], Spanish psychology students felt that animal research was a necessary means of scientific advance. In 2021, another Spanish survey sought the opinions of people who had worked with animals in research [24]. The results indicated differing ethical concerns when experimentation concerned the use of monkeys and mice compared to that of companion animals (dogs) and farm animals (pigs).

The aim of the present study was to describe current opinions and concerns about animal research, as well as knowledge of the topic among the general Spanish population.

### 2. Materials and Methods

### 2.1. Study Design

A form that did not allow duplicate responses to be given was created on the Google Forms<sup>®</sup> platform. The data collected from the replies to 21 questions were related to pop-

ulation demographics (age, gender, place of residence, education level and employment status), public opinion and knowledge regarding animal experimentation and alternative methods. The survey was launched online (on social media platforms) in 25 March 2021 and continued until 26 April 2021. The questionnaire was designed to obtain information or opinions regarding: agreement with the use of animals in research (Question 5), general knowledge regarding animal experimentation (Questions 6–9), the need for animal experiments for different purposes (Questions 10–13), different animal species as subjects of animal experiments (Questions 14–15), knowledge about alternative methods to animal models for use in research (Questions 16–17), required indications about the use of animal research in product development (Questions 18–18a), animal research related to SARS-CoV-2 (Questions 19–20) and the overall utility of the survey (Question 21).

While data were being collected for this study, there was an incident in which animal right activists published video images of what was supposedly animal abuse in a commercially run animal experimentation laboratory. In the remaining sections, this incident reported in the media is referred to as the "media incident", and some respondents completed the questionnaire immediately after this report was published. The questionnaire was adequately completed by 806 anonymous voluntary respondents. The data provided were carefully revised and prepared for subsequent statistical analysis (multivariable logistic regression model). The questionnaire is provided in the Supplementary Material.

### 2.2. Data Collection

A non-probabilistic snowball sampling approach was used to recruit participants. The weblink to the online survey was accessed via institutional websites, social networks (i.e., Facebook, Twitter and Instagram) and messaging systems, such as WhatsApp, Telegram and SMS. The only information that the respondents were given at the beginning was the title of the survey before starting, a brief explanatory document stating that it was an opinion survey on animal research and a direct link to the survey. Data were collected through the online questionnaire and accessible using smart phones, tablets and personal computers.

### 2.3. Statistical Analysis

Descriptive data from the answers are expressed as ratios for categorical variables, and as the median and interquartile range (IQR) for ordinal variables. A univariate logistic regression model was used in questions (questions 5-13, 16, 17 and 18-21) comparing different responses and demographic variables according to gender (man/woman/other), age (18 to 24 and over 24 years of age), area of residence (rural/urban), region where the respondents lived (Community of Madrid/others), scientific background (yes/no), education level (non-university/ university) and knowledge regarding the controversial media incident (questionnaire completed before/after). For questions with ordinal responses, two groups were created using the median as the cut-off point. A multivariate regression model was built from the univariate regression model. Variables returning a p < 0.100 in the univariate regression model were considered to be relevant and included in the multivariate logistic regression analysis. The final model was built using stepwise forward selection and backward elimination techniques. The significance levels set were p < 0.050 for forward selection and p < 0.100 for backward elimination. Odds ratios (ORs) were calculated along with their 95% confidence intervals (CI). All statistical tests were performed using the software package, SPSS, version 25 (IBM Corp., Armonk, NY, USA).

### 3. Results

### 3.1. Demographics

The survey was completed by 806 persons (64.3% women, 35.1% men and 0.6% not identifying with either of the above). The most represented age group was 18–24 years (58.3%). A total of 93.8% of the participants lived in an urban area (defined as more than 30,000 inhabitants and a population density greater than 100 inhabitants per km<sup>2</sup>). A total

of 65.4% lived within the Community of Madrid, and the remaining 31.7% were from elsewhere in the country.

When asked about education level, 73.0% reported they were undergraduates or had completed first-stage university studies or Master's or Doctorate studies. A total of 46.5% were enrolled in a science degree course or worked in science. Most (80.9%) had completed the survey before the controversial media incident on 10 April 2021 (see Section 2). The demographic data for the respondents are provided in Table 1.

Age (years) Gender Living area	18–24 25–29 30–39 40–65 >65 Women Men Other Urban	470 79 74 176 7 518 283 5	58.3 9.8 9.2 21.8 0.9 64.3 35.1
Gender	25–29 30–39 40–65 >65 Women Men Other	79 74 176 7 518 283	9.8 9.2 21.8 0.9 64.3
	25–29 30–39 40–65 >65 Women Men Other	74 176 7 518 283	9.2 21.8 0.9 64.3
	40–65 >65 Women Men Other	176 7 518 283	21.8 0.9 64.3
	>65 Women Men Other	7 518 283	0.9 64.3
	>65 Women Men Other	7 518 283	0.9 64.3
	Men Other	283	
	Men Other	283	
Living area	Other		35.1
Living area		5	
Living area	Urban		0.6
-	Urban		
	Ulball	756	93.8
	Rural	34	4.2
	Not defined	16	2.0
Place of residence			
	Community of Madrid	527	65.4
	Andalusia	77	9.6
	Castilla y León	51	6.3
	Extremadura	28	3.5
	Castilla-La Mancha	25	3.1
	Valencian Community	19	2.4
	Catalonia	19	2.4
		17	2.2
	Basque Country		
	Outside Spain	16	2.0
	Not defined	7	0.9
	Aragon	4	0.5
	Asturias	4	0.5
	Galicia	4	0.5
	Canary Islands	4	0.5
	Balearic Islands	2	0.2
	La Rioja	2	0.2
	Navarre	1	0.1
Education level			
Non University		217	26.9
	Non-further education (including vocational training)	108	13.4
	High school	61	7.6
	Primary school	26	3.2
	Secondary school	20	2.5
	Primary education incomplete	20	0.2
University	Timury education incomplete	589	73.1
Cruversity	University first stage	459	56.9
	University first stage Graduate University studies (Master's and Doctorate)	439 130	56.9 16.1
Scientific background	No	431	53.5
	Yes	375	46.5
Completion of questionnaire			
in relation to media incident	Before	652	80.9
	After	154	80.9 19.1

Table 1. Demographic data for survey participants (N = 806).

### 3.2. Questions

### 3.2.1. Agreement with the Use of Animals in Research (Question 5)

A total of 73.1% of the respondents stated that they agreed with animal research. Among those that were more likely to say they were opposed were women, people without a scientific background and those who had completed the questionnaire just after the media incident (OR = 1.77; 95%CI: 1.24–2.53; p = 0.002, OR = 2.47; 95%CI: 1.76–3.47; p < 0.001, OR = 2.41; 95%CI: 1.64–3.54; p < 0.001, respectively).

### 3.2.2. General Knowledge Regarding Animal Experimentation (Questions 6–9)

When the respondents were asked about how much animal research they thought was being conducted or how much information they had on it, the scores obtained were medians of 8 (IQR: 7–9) and 3 (IQR: 2–5), respectively. However, those with a scientific background and those responding before the media incident (OR = 3.12; 95%CI: 2.33–4.18; p < 0.001, OR = 1.55; 95%CI: 1.06–2.27, p = 0.023, respectively) were more likely to report if they were familiar with the topic of animal research.

When they were asked about legislation, 64.4% of the respondents claimed that they were aware of strict animal research regulations, but only 34.4% of the respondents had heard of the principle of the 3Rs by Russell and Burch. People with a scientific background (OR = 2.95; 95%CI: 2.08–4.20; p < 0.001), those who underwent university studies (OR = 2.04; 95%CI: 1.42–2.93; p < 0.001) and those who had responded before the media incident (OR = 2.50; 95%CI: 1.70–3.67; p < 0.001) were more likely to know about the existence of strict regulations. Further, respondents with a scientific background (OR = 9.43; 95%CI: 6.65–13.36; p < 0.001) showed a greater awareness of the principle of the 3Rs by Russell and Burch than the other participant subsets did.

### 3.2.3. Need for Animal Experiments for Different Purposes (Questions 10–13)

When they were asked whether animal experimentation was necessary for different purposes, a large proportion of the respondents felt it was necessary for biomedical purposes (median 8; IQR: 6–10) and to address human medical issues like diseases (median 8; IQR: 6–9). However, there was wide opposition to the use of animals to test non-medical products like cosmetics (median 2; IQR: 0–5).

Respondents who were men, older than 24 years, had a science background or had responded before the media incident (OR = 1.45; 95%CI: 1.07–1.97; p = 0.018, OR = 1.43; 95%CI: 1.04–1.96; p = 0.028, OR = 2.07; 95%CI: 1.53–2.79; p < 0.001, OR = 1.64; 95%CI: 1.12–2.40; p = 0.012, respectively) were found more likely to support animal research for biomedical purposes. Further, these subsets of participants (OR = 1.53; 95%CI: 1.13–2.02; p = 0.006, OR = 1.48; 95%CI: 1.09–2.02; p = 0.013, OR = 1.73; 95%CI: 1.28–2.32; p < 0.001, OR = 1.79; 95%CI: 1.20–2.68; p = 0.005, respectively) were also more likely to be supportive of animal research to address human diseases. Additionally, men emerged as more likely to support animal testing of non-medical products than the other groups were (OR = 1.63; 95%CI: 1.22–2.19; p = 0.001).

A total of 84.2% of respondents agreed with the use of a drug tested in the same species as their pet. Non-residents of the Community of Madrid, those with a scientific background, those with university degrees, those who responded before the media incident (OR = 1.96; 95%CI: 1.22–3.17; p = 0.006, OR = 1.88; 95%CI: 1.19–2.96; p = 0.007, OR = 1.60; 95%CI: 1.02–2.50; p = 0.042, OR = 2.03; 95%CI: 1.30–3.18; p = 0.002, respectively) or those who had a sick pet were also more likely than the remaining groups were to agree with testing medications in species for which the drug was intended.

### 3.2.4. Different Animal Species as Subjects of Animal Experiments (Questions 14–15)

More than three-quarters of the respondents reported that the animals most frequently used in animal research were mice (83.0%) and rats (78.5%). Only 26.6% of participants mentioned that primates were among the most used animals. Cats and fish were described as the least used ones (1% and 0.7%, respectively).

When asked which animals they felt should be used for research, 73.9% indicated they would choose mice, followed by flies, non-human primates and fish (32.6%, 26.1% and 16.6%, respectively). Only 15.1% were indifferent to the use of a given animal species for research.

# 3.2.5. Knowledge about Alternative Methods to Animal Models for Use in Research (Questions 16 and 17)

When they were asked if they knew about alternatives to the use of animals in research, 77.8% of respondents stated there were alternatives. Compared to other demographic groups, women, those living in urban areas and those with a scientific background (OR = 1.48; 95%CI: 1.04–2.11; p = 0.030, OR = 3.40; 95%CI: 1.65–6.99; p = 0.001, OR = 2.94; 95%CI: 2.01–4.30; p < 0.001, respectively) were much more likely to know about the existence of alternative methods. Respondents who mentioned this were asked to describe the methods they knew about. The best-known methods were cell lines (58.4%), followed by invertebrates (22.0%), in silico models (20.9%) and organ microchip models (19.9%).

Further, when asked about this topic, 73.9% of respondents agreed that these alternative methods should be employed more often than animal models are. Women and those without a scientific background (OR = 1.79; 95%CI: 1.29–2.48; p < 0.001, OR = 1.40; 95%CI: 1.02–1.93; p = 0.04, respectively) were more likely agree with this idea.

3.2.6. Require Indications about the Use of Animal Research in Product Development (Questions 18 and 18a)

The vast majority (87.2%) of participants stressed that all commercial products should clearly state whether they have been tested on animals. Of all respondents, 50.5% said they would stop using products if they did not specify whether they had been tested on animals. Women (OR = 2.71; 95%CI: 1.78–4.13; p < 0.001) were more likely to differ from other groups in their belief that all products must have an indication of whether they have been subjected to animal testing. In addition, the participant subsets women (OR = 2.08; 95%CI: 1.54–2.81; p < 0.001), those without a scientific background (OR = 1.41; 95%CI: 1.06–1.89; p = 0.018) and people who responded after the media incident (OR = 2.01; 95%CI: 1.38–2.92; p < 0.001) were more likely to state they would not continue to use a product that has been since tested on animals.

### 3.2.7. Animal Research Related to SARS-CoV-2 (Questions 19-20)

Animal research was deemed to be essential for the study of SARS-CoV-2 by 82.3% of respondents, although 28.9% agreed it should only be used to address medical emergencies such as the COVID-19 pandemic. According to a majority of men, participants with a scientific background and those responding before the media incident, animal research was probably crucial to understanding SARS-CoV-2 and developing treatments (OR = 2.21; 95%CI: 1.44–3.41; p < 0.001, OR = 2.04; 95%CI: 1.39–3.00; p < 0.001, OR = 1.79; 95%CI: 1.17–2.76; p = 0.008, respectively). Individuals without a scientific background and people who completed the survey after the media event (OR = 3.27; 95%CI: 2.34–4.58; p < 0.001, OR = 1.92; 95%CI: 1.31–2.83; p = 0.001, respectively) were more likely to state that animal testing should be reserved for resolving critical issues.

### 3.2.8. Overall Utility of the Survey (Question 21)

Among the respondents, 82.6% said that the survey had made them think about animal research. Women (OR = 2.06; 95%CI: 1.41–3.02; p < 0.001), respondents without a scientific background (OR = 1.89; 95%CI: 1.29–2.76; p = 0.001) and those who completed the questionnaire after the media incident (OR = 2.85; 95%CI: 1.52–5.33; p = 0.001) were more likely to say they were concerned about this.

In Tables 2 and 3, we present all the information regarding the survey responses and their analyses, respectively. The results of our univariate test and the non-significant variables for the multivariate test are detailed in Table S1 provided in the Supplementary Material.

# **Table 2.** Survey responses (Questions 5–21).

Variables		n (N = 806)	%
Do you approve of using animals for research purposes? (Question 5)			
	Yes	589	73.1
	No	217	26.9
How much animal research do you think is done			
(from 0 to 10, where $0 = \text{none and } 10 = a \text{ lot}$ )? (Question 6)			
		8.0 *	(7.0–9.0) **
How much do you think you know about animal research e.g., protocols,			
legislation, etc (from 0–10, where $0 =$ nothing and $10 =$ a lot)? (Question 7)			
		3.0 *	(2.0-5.0) **
Do you think that animal research is subject to strict regulation? (Question 8)			
	No	287	35.6
	Yes	519	64.4
Do you know the principle of the 3Rs by Russell and Burch? (Question 9)			
	No	529	65.6
	Yes	277	34.4
How necessary do you think animal experimentation is for biomedical			
research (from $0-10$ , where $0 = $ not at all and $10 = $ very)? (Question 10)			
······································		8.0 *	(6.0-10.0) **
How much do you agree with using animals to find a cure for human			. /
diseases (from 0–10, where 0 = not at all and 10 = very much)? (Question 11)			
$\frac{1}{1} = \frac{1}{1} + \frac{1}$		8.0 *	(6.0–9.0) **
			(0.0 7.0)
If you have or had a sick pet that needed medication, would you agree that this medication should be tested on the same species as your pet for a drug			
this medication should be tested on the same species as your pet for a drug agency to approve it? (Question 12)			
agency to approve it. (Question 12)	No	127	15.8
	Yes	679	84.2
How much do you correct with using animals to test non-medicinal products			
How much do you agree with using animals to test non-medicinal products (from $0-10$ , where $0 = not$ at all and $10 = very much$ )? (Question 13)			
(from 0 10, where 0 = not at an and 10 = very mach). (Question 10)		2.0 *	(0.0-5.0) **
		2.0	(010 010)
Which animals do you think are the most used in research? (Question 14)	Na	127	17.0
Mouse	No Yes	137 669	17.0 83.0
Rat	No	173	21.5
Rat	Yes	633	78.5
Non-human primate	No	594	73.7
- ···· I ······	Yes	212	26.3
Dog	No	772	95.8
0	Yes	34	4.2
Rabbit	No	785	97.4
	Yes	21	2.6
Invertebrates	No	790	98.0
	Yes	16	2.0
Farm animals	No	793	98.4
	Yes	13	1.6
Cat	No Voc	794	98.5 1 5
Others	Yes No	12 798	1.5 99.0
Ulicity	No Yes	798	99.0 1.0
Fish	No	800	99.3
	Yes	6	0.7
$M_{\rm bish} animal would not share (-1) (0, 1) (1) (1) (1) (1) (1) (1) (1) (1) (1) $		~	•
Which animal would you choose for biomedical research? (Question 15) Mouse	No	210	26.1
wouse	No Yes	210 596	26.1 73.9
Drosophila melanogaster (fruit fly)	No	543	67.4
2. coopting montograde (it the ity)	Yes	263	32.6
Non-human primate	No	596	73.9
I	Yes	210	26.1
Fish	No	672	83.4
	Yes	134	16.6
Indifferent	No	684	84.9
	Yes	122	15.1
		777	06.4
Dog	No Yes	777 29	96.4 3.6

# Table 2. Cont.

Variables		n (N = 806)	%
Do you think there are alternative methods to animal models for research? (Question 16)			
	No	179	22.2
	Yes	627	77.8
If you think there are alternative methods to animal models for research, indicate the ones you know about. (Question 16a)			
In vitro cell lines	No	261	41.6
	Yes	366	58.4
Invertebrate animal models	No	489	78.0
	Yes	138	22.0
In silico analysis	No	496	79.1
	Yes	131	20.9
Tissue models: chips	No	508	81,0
houe mouels, empo	Yes	119	19.0
Do you think these alternative methods should be used more than animal models? (Question 17)			
mould. (Question II)	No	210	26.1
	Yes	596	73.9
Do you think it should be mandatory for all products to state whether animal testing was necessary for their preparation? (Question 18)			
	No	103	12.8
	Yes	703	87.2
If you answered yes, would you stop using a product that previously did not provide that information but now states it was teste in animals ? (Question 18a)			
	No	399	49.5
	Yes	407	50.5
Do you think that animal research has been or is still necessary to find a cure for COVID-19 (e.g., vaccine, treatment)? (Question 19)			
	No	143	17.7
	Yes	663	82.3
Do you think that scientists should conduct animal testing only in certain situations such as the SARS-CoV-2 emergency, and not for other diseases? (Question 20)			
(Zuesuon 20)	No	573	71.1
	Yes	233	28.9
Has this survey made you think about animal research? (Question 21)			
This this survey made you think about animal research: (Question 21)	No	140	17.4
	Yes	666	82.6

\* Data expressed as median. \*\* Data expressed as IQR.

# **Table 3.** Multivariate regression model of Questions from 5 to 13 and from 16 to 21.

Question 5. Do you approve of us	ing animals for research	h purposes?						
Variable		Participation (n)						
		Yes	No	OR	95%CI			<i>p</i> -Value
Gender	Men	225	58			1.00		
	Women	362	156	1.77	1.24	-	2.53	0.002
Scientific background	Yes	304	71			1.00		
Ū	No	285	146	2.47	1.76	-	3.47	< 0.001
Media incident	Before	499	153			1.00		
	After	90	64	2.41	1.64	-	3.54	< 0.001
Question 6. How much animal res	earch do you think is d	lone (0 = none to 10 = a	lot)?					
¥7 · 11	Participation (n)			Multivariate				
Variable		<median (<8.0)<="" td=""><td><math>\geq</math>median (<math>\geq</math>8.0)</td><td>OR</td><td></td><td>95%CI</td><td></td><td><i>p</i>-Value</td></median>	$\geq$ median ( $\geq$ 8.0)	OR		95%CI		<i>p</i> -Value
Conden	Men	119	164			1.00		
Gender	Women	181	337	1.35	1.00	-	1.82	0.047

		Participa	ation (n)			Multivariate	5	
Variable		≤median (≤3.0) >median (>3.0)		OR	95%CI			<i>p</i> -Value
Scientific background	No	302	129			1.00		r
Serenine Suchground	Yes	163	212	3.12	2.33	-	4.18	< 0.001
Media incident	After	99	55			1.00		
	Before	366	286	1.55	1.06	-	2.27	0.023
Question 8. Do you think that an	imal research is subject to	strict regulation?						
X7 · 11		Participa	ation ( <i>n</i> )		Multivariate			
Variable		No	Yes	OR		95%CI		<i>p</i> -Value
Scientific background	No	209	222			1.00		•
Scientific Duckground	Yes	78	297	2.95	2.08	-	4.20	< 0.001
Educational level university	No	122	95			1.00		
	Yes	165	424	2.04	1.42	-	2.93	< 0.001
Media incident	After	78	76			1.00		
	Before	209	443	2.50	1.70	-	3.67	< 0.001
Question 9. Do you know the pri	nciple of the 3Rs by Russ	ell and Burch?						
Variable		Participa			Multivariate	5		
Vallable		No	Yes	OR		95%CI		<i>p</i> -Value
Scientific background	No	375	56			1.00		
0	Yes	154	221	9.43	6.65	-	13.36	< 0.001
Question 10. How necessary do y	ou think animal experime	entation is for biomed	ical research (0 = not	at all to 10	= very)?			
		Participa	ation ( <i>n</i> )			Multivariate	2	
Variable		<median (<8.0)<="" td=""><td>&gt;median (&gt;8.0)</td><td>OR</td><td></td><td>95%CI</td><td></td><td>p-Value</td></median>	>median (>8.0)	OR		95%CI		p-Value
Ago	18–24	215	255			1.00		,
Age	>24	126	233	1.43	1.04	1.00	1.96	0.028
Gender	Women	233	285	1.45	1.04	1.00	1.70	0.020
	Men	105	178	1.45	1.07	-	1.97	0.018
Scientific background	No	211	220			1.00		
	Yes	130	245	2.07	1.53	-	2.79	< 0.001
Media incident	After Before	84 257	70 395	1.64	1.12	1.00	2 40	0.012
						-	2.40	0.012
Question 11. How much do you a	igree with using animals t			all to $10 =$	very much			
Variable		Participa				Multivariate	2	
		$\leq$ median ( $\leq$ 8.0)	>median (>8.0)	OR		95%CI		<i>p</i> -Value
Age	18-24	290	180	1.40	1.00	1.00		0.010
Carlan	>24	171	165 205	1.48	1.09	1.00	2.02	0.013
Gender	Women Men	313 143	203 140	1.53	1.13	-	2.02	0.006
Scientific background	No	265	140	1.00	1.15	1.00	2.02	0.000
Selentine Suchground	Yes	196	179	1.73	1.28	-	2.32	< 0.001
Media incident	After	109	45			1.00		
	Before	352	300	1.79	1.2	-	2.68	0.005
Question 12. If you have or had a lrug agency to approve it?	sick pet that needed med	ication, would you ag	ree that this medicati	on should	oe tested or	n the same spe	cies as you	r pet for a
ing agency to approve it:		Participa	ation (n)			Multivariate		
Variable		No	Yes	OR		95%CI	-	<i>p</i> -Value
	NT			OK				<i>p</i> -value
Scientific background	No Yes	88 39	343 336	1.88	1.19	1.00	2.96	0.007
Autonomous Community	Madrid	100	427	1.00	1.17	1.00	2.70	0.007
······································	Other communities	25	229	1.96	1.22	-	3.17	0.006
Educational level university	No	51	166			1.00		
	Yes	76	513	1.60	1.02	-	2.50	0.042
Media incident	After Before	38 89	116 563	2.03	1.30	1.00	3 10	0.002
Numetica 10 Hos						-	3.18	0.002
Question 13. How much do you a	gree with using animals l	-		1  to  10 = ve	ry much)?	<b>X</b> 10		
Variable		Participa				Multivariate	2	
		$\leq$ median ( $\leq$ 2.0)	>median (>2.0)	OR		95%CI		<i>p</i> -Value
Conton	Women	301	217			1.00		
Gender	Men	130	153	1.63	1.22		2.19	0.001

### Table 3. Cont.

		Particin	ation (n)			Multivariate		
Variable		<b>`</b>		OR		95%CI		u Valua
		No		UK				<i>p</i> -Value
Gender	Men	80	203	1.10	1.04	1.00	0.11	0.00
	Women	99	419	1.48	1.04	-	2.11	0.03
Place of residence	Rural	17	17 598	2 40	1.65	1.00	6.99	0.001
Scientific background	Urban No	158 133	598 298	3.40	1.65	1.00	6.99	0.001
Scientific Dackground	Yes	46	329	2.94	2.01	1.00	4.30	< 0.001
Owned and 17 De serve de sale de serve al				2.74	2.01		4.50	<0.001
Question 17. Do you think these al	ternative methods shot					Multivariate		
Variable		Particip	oation (n)					
		No	Yes	OR		95%CI		<i>p</i> -Value
Gender	Men	94	189			1.00		
	Women	116	402	1.79	1.29	-	2.48	< 0.001
Scientific background	Yes	108	267			1.00		
	No	102	329	1.40	1.02	-	1.93	0.04
Question 18. Do you think it shoul	d be mandatory for all	products to state whet	her animal testing	was necessary	for their pi	reparation?		
¥7 · 11		Particip	ation (n)			Multivariate		
Variable	Variable		Yes	OR		95%CI		<i>p</i> -Value
Gender	Men	58	225			1.00		
Genuer	Women	45	473	2.71	1.78	-	4.13	< 0.001
Question 18a. If you answered yes,	would you stop using	a product that previou	sly did not provid	e that informa	tion but nov	w states it was	tested in a	nimals?
~,	, i i j i i i i i i i i i i i i i i i i		pation (n)			Multivariate		
Variable		No	Yes	OR		95%CI		<i>p</i> -Value
				OK				<i>p</i> -value
Gender	Men	173	110			1.00		
	Women	224	294	2.08	1.54	-	2.81	< 0.001
Scientific background	Yes	198	177		1.07	1.00	1.00	0.010
No. 11. 1. 1. 1. 1.	No	201	230	1.41	1.06	-	1.89	0.018
Media incident	Before After	345 54	307 100	2.01	1.38	1.00	2.92	< 0.001
Question 10 Demonsthinds that and							2.72	<0.001
Question 19. Do you think that ani	mai research has been o			/ID-19 (e.g., va	accine, treat			
Variable			Participation ( <i>n</i> )			Multivariate		
		No	Yes	OR		95%CI		<i>p</i> -Value
Gender	Women	110	408			1.00		
	Men	32	251	2.21	1.44	-	3.41	< 0.001
Scientific background	No	94	337			1.00		
	Yes	49	326	2.04	1.39	-	3.00	< 0.001
Media incident	After	40	114			1.00		
	Before	103	549	1.79	1.17	-	2.76	0.008
		minual testine only in	certain situations s	such as the SA	RS-CoV-2 e	mergency, and	not for otl	er diseases
Question 20. Do you think that scie	entists should conduct a	inimal testing only in				Multivariate		
· ·	entists should conduct a		pation (n)			withivallate		87.1
Question 20. Do you think that scie Variable	entists should conduct a			OR		95%CI		<i>p</i> -Value
· ·	Pentists should conduct a	Particip	eation ( <i>n</i> ) Yes 64	OR				1
Variable Scientific background	Yes No	Particip No 311 262	Pation (n) Yes 64 169	OR 3.27	2.34	95%CI	4.58	<i>p</i> -value <0.001
Variable	Yes No Before	Particip No 311 262 478	64 169 174	3.27		95%CI		<0.001
Variable Scientific background	Yes No	Particip No 311 262	Pation (n) Yes 64 169		2.34 1.31	95%CI	4.58 2.83	1
Variable Scientific background	Yes No Before After	Particip No 311 262 478 95	64 169 174	3.27		95%CI		<0.001
Variable Scientific background Media incident Question 21. Has this survey made	Yes No Before After	Particip No 311 262 478 95 l research?	64 169 174	3.27		95%CI	2.83	<0.001
Variable Scientific background Media incident	Yes No Before After	Particip No 311 262 478 95 l research?	eation (n) Yes 64 169 174 59	3.27		95%CI 1.00 - 1.00 -	2.83	<0.001
Variable Scientific background Media incident Question 21. Has this survey made Variable	Yes No Before After <b>you think about anim</b> a	Particip No 311 262 478 95 I research? Particip No	Pation (n) Yes 64 169 174 59 ration (n) Yes	3.27 1.92		95%CI 1.00 - 1.00 - Multivariate 95%CI	2.83	<0.001 0.001
Variable Scientific background Media incident Question 21. Has this survey made	Yes No Before After <b>you think about anima</b> Men	Particip           No           311           262           478           95           I research?           Particip           No           68	ration (n) Yes 64 169 174 59 ration (n) Yes 215	3.27 1.92 OR	1.31	95%CI 1.00 - 1.00 - Multivariate	2.83	<0.001 0.001 <i>p</i> -Value
Variable Scientific background Media incident Question 21. Has this survey made Variable Gender	Yes No Before After <b>you think about anim</b> a	Particip           No           311           262           478           95           I research?           Particip           No           68           71	ration (n) Yes 64 169 174 59 ration (n) Yes 215 447	3.27 1.92		95%CI 1.00 - 1.00 - Multivariate 95%CI 1.00 -	2.83	<0.001 0.001
Variable Scientific background Media incident Question 21. Has this survey made Variable	Yes No Before After <b>you think about anima</b> Men Women Yes	Particip No 311 262 478 95 l research? Particip No 68 71 80	ration (n) Yes 64 169 174 59 ration (n) Yes 215 447 295	3.27 1.92 OR 2.06	1.31	95%CI 1.00 - 1.00 - Multivariate 95%CI	2.83	<0.001 0.001 <i>p</i> -Value <0.001
Variable Scientific background Media incident Question 21. Has this survey made Variable Gender	Yes No Before After <b>you think about anima</b> Men Women	Particip           No           311           262           478           95           I research?           Particip           No           68           71	ration (n) Yes 64 169 174 59 ration (n) Yes 215 447	3.27 1.92 OR	1.31	95%CI 1.00 - 1.00 - Multivariate 95%CI 1.00 -	2.83	<0.001 0.001 <i>p</i> -Value

### Table 3. Cont.

## 4. Discussion

Animal research generates controversy due to bioethical concerns [27,28]. This survey was designed to assess opinions about animal research in Spain. The questionnaire was completed by 806 persons. The survey tried to consider the opinions of different strata of society in Spain.

The questionnaire was designed to cover all necessary aspects when there is no specific target population. However, the main limitation of our study was that, by conducting the survey online, it is probably that fifty percent the respondents were aged 18–24 years and that only a small percentage were over 65 years. This could make the results non-extrapolatable to other age groups. We should also mention that while the initial population was largely made up of young university students living in Madrid, by making the questionnaire easily accessible to their close contacts via social media, we were able to reach other sectors of the Spanish population. Further research on this topic should ensure a targeted distribution of the questionnaire with a structural design to ensure a proper representation of the population and avoid a skewed distribution by gender, age, academic background, or even survey distribution route with an over-representation of certain sectors of the population that may limit the applicability of the results to the wider population.

Our findings revealed that over 70% of the respondents support the use of animals for research. This figure is slightly higher than that described for a given region of our country by Navarro et al. [26] in 2001, in that 65.7% of respondents strongly agreed or agreed with animal research; women and those with a non-scientific background were more likely to be against animal experimentation. Other authors also detected gender differences in the attitudes toward animal research. These differences have been explained by gender variations in socialization, attributing an emphasis on caring, nurturing and expressiveness to women rather than men [29–32]. In these studies, it was also concluded that men emphasized more the potential benefits arising from the use of animals in research [29–32]. It also seems that the general public's lack of knowledge of the topic makes persons more sensitive to animals' rights and suffering, leading them to question the real benefits of animal experimentation, as has been shown in other studies [33]. However, other authors suggest that these attitudes are less about a lack of knowledge and more about a perspective that comes with particular lifestyles, viewpoints and access to types of media [34]. In fact, it has been argued that as the level of awareness increases, the public may become less supportive, especially if the issue under discussion is considered to be morally contentious [35]. In contrast, people with a scientific background are likely to have a more informed opinion because of the scientific nature of the topic [32,36]. From a science communication perspective, this assumes that science is inherently the best way to acquire knowledge, and for most of the population, that is not a given fact. However, the tendency of scientifically literate people to adopt a more utilitarian perspective can, among other factors, mean that they see the necessity of using animals in research [37].

Similar to the UK study [28], our study reveals there is minimal knowledge about animal welfare regulations and animal care and handling guidelines among the Spanish public. However, it was also observed here that people feel that animal research is needed for developments in the area of human health and safety, but not for other purposes, such as testing cosmetic products. In effect, the respondents of a survey by Uchoshiki et al. [27] in 2019 reported that they approved of animal experimentation as long as it was for biomedical, and not cosmetic, purposes. In our study, a large percentage of the respondents mentioned that animal research is subject to strict regulation. This attitude was probably more common among those with a scientific background and higher education level. As seen in previous studies, having a scientific qualification and training means that respondents tend to assume they are more knowledgeable about animal research [32,36]. In Spain, animal research is regulated by Royal Decree 53/2013 [38] and the recommendations of the European Union provided in EU Directive 2010/63 [13]. These regulations are based on the principle of the 3Rs by Russell and Burch. The 3Rs are unknown to a large proportion of the general population, which focuses on the welfare of experimental animals [36], as an issue of great public concern [39]. This was also observed in the study by Iki et al. in 2017 [40], indicating a growing awareness of animal bioethics. This percentage is slightly higher than those in previous reports. This finding could be related to the characteristics of our population, with an important representation of respondents have a scientific background.

Our respondents preferred the use of mice (Mus musculus) for biomedical research above other species. This is consistent with the results of a Spanish survey performed in 2021, in which it was found that a higher percentage of people were against the use of dogs compared to those against the use of mice [24].

Motivated by the principle of the 3Rs, alternative methods have started to replace the use of animals whenever possible, and today, several of these alternative methods have roles in research [23]. Indeed, our survey showed that these models were known to most respondents. In another study [28], support for animal experimentation was linked to a lack of alternatives. This indicates respondents' awareness of the use of alternative methods and was, in fact, evident in our survey, as a large percentage of respondents (73.9%) felt that alternatives should be used instead of experimental animals.

Another important factor that affected our results was that, at the time of our survey, an animal welfare NGO released video images reportedly taken at a pharmaceutical/biotechnological research company in Madrid [41]. The video recording depicted practices allegedly fulfilling established regulations involving verbal insults to animals and mishandling. It also showed questionable procedures performed on beagles, monkeys, mice, rabbits and pigs. That event caused an opinion bias in our respondents, in that almost twice as many people were against animal research when compared to the data obtained from forms completed earlier. As a result, animal research regulations and research for human biomedical purposes were questioned, as the video images did not seem to comply with animal welfare standards [13,38]. Despite this change in opinion, alternative methods, which are widely used in toxicological tests, have not been publicized [23]. As previously described [35], misinformation, erroneous information and the propagation of information through social networks produce alarm in some strata of society, and this leads to changing opinions or subjective opinions that are far removed from reality. It is important to emphasize that our survey indicated that the people who mentioned they had thought most about the questionnaire were those most likely to be against animal experimentation.

This study shows that, despite the fact that a large percentage of the population supports the use of research animals, it constitutes a sensitive issue about which there is a certain lack of knowledge, and it can be manifestly altered by events in a short period of time. In this sense, it is important that those responsible for formulating policies to improve those measures aimed at the transparency of the use of animals through awareness programs on this matter so that they receive impartial, unbiased information. In the same way, research facilities must always apply, in accordance with the law, measures that guarantee the animal welfare [42–44].

### 5. Conclusions

Our findings indicate that animal research is a sensitive social issue in Spain, mainly involving ethical and moral concerns about the use of animals in research. These concerns were sometimes motivated by a lack of information on this topic. Opinions seemed to be conditioned by gender and education level. It was also observed here that events reported in the media may cause sudden changes in public opinion. This study has highlighted the lack of information on animal experimentation provided by the scientific community to society. A potential solution to this knowledge gap could involve the integration of educational programs in countries, aimed at fostering positive attitudes and providing training to future generations regarding animal research.

**Supplementary Materials:** The following supporting information can be downloaded at: https://www.mdpi.com/article/10.3390/ani13122039/s1, Survey; Table S1. Univariate and multivariate regression model of questions 5 to 13 and 16 to 21.

**Author Contributions:** A.M.-B. wrote the manuscript, contributed to the design of the survey, data analysis and she also contributed to preparing the tables. M.F.-R. contributed to the conception and design of the manuscript, data analysis, preparing the tables, drafting the manuscript, and reviewed successive drafts of the paper. D.D.-R. contributed to the design of the survey, drafting the manuscript, and reviewed successive drafts of the paper. G.O.-D. contributed to the design of the statistical analysis, data analysis, preparing the tables and reviewed the manuscript. L.R. contributed to the conception and design of the survey and funding acquisition. J.A.D.P.-M. contributed to the conception and design of the survey and the manuscript, data analysis, drafting the manuscript, preparing the tables, and reviewed successive drafts of the paper. J.A.D.P.-M. is the principal investigator. All authors have read and agreed to the published version of the manuscript.

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**Institutional Review Board Statement:** The questionnaire was anonymous, public, and freely accessible and had no special incentives targeting only certain adult members of the public. No personal information was collected from respondents such as name, address, telephone number, or e-mail address. In accordance with Regulation (EU) 2016/679 of the European Parliament and of the Council of 27 April 2016, and Organic Law 3/2018, of 5 December, on the Protection of Personal Data and Guarantee of Digital Rights, this study is not obligated to have an ethics committee since no personal data were used and the survey was conducted anonymously. Additionally, it was clearly stated in the survey's preamble that the data would be utilized solely for scientific purposes. As no animals were used in this study, the need for ethics approval was waived.

Informed Consent Statement: Informed consent was obtained from all subjects involved in the study.

**Data Availability Statement:** The original contributions are included in the article/Supplementary Material, further inquiries can be directed to the corresponding author.

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## References

- 1. Petetta, F.; Ciccocioppo, R. Public perception of laboratory animal testing: Historical, philosophical, and ethical view. *Addict. Biol.* **2021**, *26*, e12991. [CrossRef]
- Rahman, M.M.; Ghoshal, U.C.; Ragunath, K.; Jenkins, G.; Rahman, M.; Edwards, C.; Hasan, M.; Taylor-Robinson, S.D. Biomedical research in developing countries: Opportunities, methods, and challenges. *Indian J. Gastroenterol.* 2020, 39, 292–302. [CrossRef]
- 3. Erick Peirson, B.R.; Kropp, H.; Damerow, J.; Laubichler, M.D. The diversity of experimental organisms in biomedical research may be influenced by biomedical funding. *Bioessays* 2017, *39*, 1600258. [CrossRef] [PubMed]
- 4. Polejaeva, I.A.; Rutigliano, H.M.; Wells, K.D. Livestock in biomedical research: History, current status and future prospective. *Reprod. Fertil. Dev.* **2016**, *28*, 112–124. [CrossRef] [PubMed]
- 5. Ericsson, A.C.; Crim, M.J.; Franklin, C.L. A brief history of animal modeling. Mol. Med. 2013, 110, 201–205.
- 6. Meigs, L.; Smirnova, L.; Rovida, C.; Leist, M.; Hartung, T. Animal testing and its alternatives—The most important omics is economics. *ALTEX* **2018**, *35*, 275–305. [CrossRef] [PubMed]
- 7. Robinson, N.B.; Krieger, K.; Khan, F.M.; Huffman, W.; Chang, M.; Naik, A.; Yongle, R.; Hameed, I.; Krieger, K.; Girardi, L.N.; et al. The current state of animal models in research: A review. *Int. J. Surg.* **2019**, *7*2, 9–13. [CrossRef]
- 8. Sun, S.H.; Chen, Q.; Gu, H.J.; Yang, G.; Wang, Y.X.; Huang, X.Y.; Liu, S.S.; Zhang, N.N.; Li, X.F.; Xiong, R.; et al. A Mouse Model of SARS-CoV-2 Infection and Pathogenesis. *Cell Host Microbe* **2020**, *28*, 124–133.e4. [CrossRef]
- Rahman, M.T.; Sobur, M.A.; Islam, M.S.; Ievy, S.; Hossain, M.J.; El Zowalaty, M.E.; Rahman, A.T.; Ashour, H.M. Zoonotic Diseases: Etiology, Impact, and Control. *Microorganisms* 2020, *8*, 1405. [CrossRef]
- Orusa, T.; Orusa, R.; Viani, A.; Carella, E.; Borgogno Mondino, E. Geomatics and EO Data to Support Wildlife Diseases Assessment at Landscape Level: A Pilot Experience to Map Infectious Keratoconjunctivitis in Chamois and Phenological Trends in Aosta Valley (NW Italy). *Remote Sens.* 2020, 12, 3542. [CrossRef]
- Viani, A.; Orusa, T.; Borgogno-Mondino, E.; Orusa, R. Snow Metrics as Proxy to Assess Sarcoptic Mange in Wild Boar: Preliminary Results in Aosta Valley (Italy). *Life* 2023, 13, 987. [CrossRef] [PubMed]
- 12. Akhtar, A. The flaws and human harms of animal experimentation. *Camb. Q. Healthc. Ethics* **2015**, *24*, 407–419. [CrossRef] [PubMed]
- European Parliament and the Council. Directive 2010/63/EU: Protection of Animals Used for Scientific Purposes. 2010. Available online: https://eur-lex.europa.eu/legal-content/EN/ALL/?uri=CELEX%3A32010L0063 (accessed on 11 May 2023).

- 14. US Government. The Animal Welfare Act (AWA). 1990. Available online: https://www.nal.usda.gov/animal-health-and-welfare/animal-welfare-act (accessed on 11 May 2023).
- 15. UK Government. Animal Welfare Act. 2006. Available online: https://www.legislation.gov.uk/ukpga/2006/45/contents (accessed on 11 May 2023).
- 16. Cheluvappa, R.; Scowen, P.; Eri, R. Ethics of animal research in human disease remediation, its institutional teaching; and alternatives to animal experimentation. *Pharmacol. Res. Perspect.* **2017**, *5*, e00332. [CrossRef] [PubMed]
- 17. Hartung, T.; Blaauboer, B.; Leist, M. Food for thought ... on education in alternative methods in toxicology. *ALTEX* 2009, 26, 255–263. [CrossRef]
- Daneshian, M.; Akbarsha, M.A.; Blaauboer, B.; Caloni, F.; Cosson, P.; Curren, R.; Goldberg, A.; Gruber, F.; Ohl, F.; Pfaller, W.; et al. A framework program for the teaching of alternative methods (replacement, reduction, refinement) to animal experimentation. *ALTEX* 2011, 28, 341–352. [CrossRef]
- 19. Bracken, M.B. Why animal studies are often poor predictors of human reactions to exposure. *J. R. Soc. Med.* **2009**, *102*, 120–122. [CrossRef]
- Khoury, D.S.; Wheatley, A.K.; Ramuta, M.D.; Reynaldi, A.; Cromer, D.; Subbarao, K.; O'Connor, D.H.; Kent, S.J.; Davenport, M.P. Measuring immunity to SARS-CoV-2 infection: Comparing assays and animal models. *Nat. Rev. Immunol.* 2020, 20, 727–738. [CrossRef]
- Ribitsch, I.; Baptista, P.M.; Lange-Consiglio, A.; Melotti, L.; Patruno, M.; Jenner, F.; Schnabl-Feichter, E.; Dutton, L.C.; Connolly, D.J.; van Steenbeek, F.G.; et al. Large Animal Models in Regenerative Medicine and Tissue Engineering: To Do or Not to Do. Front. *Bioeng. Biotechnol.* 2020, *8*, 972. [CrossRef]
- 22. Swearengen, J.R. Choosing the right animal model for infectious disease research. *Anim. Models Exp. Med.* **2018**, *1*, 100–108. [CrossRef]
- 23. Freires, I.A.; Sardi, J.C.; de Castro, R.D.; Rosalen, P.L. Alternative Animal and Non-Animal Models for Drug Discovery and Development: Bonus or Burden? *Pharm. Res.* 2017, *34*, 681–686. [CrossRef]
- 24. Goñi-Balentziaga, O.; Ortega-Saez, I.; Vila, S.; Azkona, G. A survey on the use of mice, pigs, dogs and monkeys as animal models in biomedical research in Spain. *Lab. Anim. Res.* **2022**, *38*, 14. [CrossRef] [PubMed]
- 25. Merkley, R.; Pippin, J.J.; Joffe, A.R. A survey to understand public opinion regarding animal use in medical training. *Altern. Lab. Anim.* **2018**, *46*, 133–143. [CrossRef] [PubMed]
- 26. Navarro, J.F.; Maldonado, E.; Pedraza, C.; Cavas, M. Attitudes toward animal research among psychology students in Spain. *Psychol. Rep.* **2001**, *89*, 227–236. [CrossRef] [PubMed]
- 27. Uchikoshi, A.; Kasai, N. Survey report on public awareness concerning the use of animals in scientific research in Japan. *Exp. Anim.* **2019**, *68*, 307–318. [CrossRef]
- Clemence, M.; Leaman, J. Public Attitudes to Animal Research in 2016. 2016. Available online: https://doc.ukdataservice.ac.uk/ doc/8059/mrdoc/pdf/8059\_ols\_public\_attitudes\_to\_animal\_research\_report.pdf (accessed on 11 May 2023).
- 29. Eldridge, J.J.; Gluck, J.P. Gender differences in attitudes toward animal research. Ethics Behav. 1996, 6, 239–256. [CrossRef]
- 30. Hagelin, J.; Carlsson, H.-E.; Hau, J. An overview of surveys on how people view animal experimentation: Some factors that may influence the outcome. *Public Underst. Sci.* 2003, 12, 67–81. [CrossRef]
- 31. Herzog, H.A.; Betchart, N.S.; Pittman, R.B. Gender, Sex Role Orientation, and Attitudes toward Animals. *Anthrozoös* **1991**, *4*, 184–191. [CrossRef]
- 32. Sandgren, E.P.; Streiffer, R.; Dykema, J.; Assad, N.; Moberg, J. Attitudes toward animals, and how species and purpose affect animal research justifiability, among undergraduate students and faculty. *PLoS ONE* **2020**, *15*, e0233204. [CrossRef]
- 33. Ormandy, E.H.; Schuppli, C.A. Public Attitudes toward Animal Research: A Review. Animals 2014, 4, 391–408. [CrossRef]
- 34. Riddle, E.; MacKay, J.R.D. Social Media Contexts Moderate Perceptions of Animals. Animals 2020, 10, 845. [CrossRef]
- 35. van der Linden, S. Misinformation: Susceptibility, spread, and interventions to immunize the public. *Nat. Med.* **2022**, *28*, 460–467. [CrossRef] [PubMed]
- Hagelin, J.; Hau, J.; Carlsson, H.E. Attitude of Swedish veterinary and medical students to animal experimentation. *Vet. Rec.* 2000, 146, 757–760. [CrossRef] [PubMed]
- 37. Reddy, D. Scientific literacy, public engagement and responsibility in science. Cult. Sci. 2021, 4, 6–16. [CrossRef]
- Government of Spain. Royal Decree 53/2013: Establishing the Basic Rules Applicable to the Protection of Animals Used in Experimentation and Other Scientific Purposes, including Teaching. 2013. Available online: <a href="https://www.boe.es/diario\_boe/txt">https://www.boe.es/diario\_boe/txt</a>. php?id=BOE-A-2013-1337 (accessed on 11 May 2023).
- 39. Festing, S. On the necessity for animal experimentation. *Bioessays* 2008, 30, 94–95. [CrossRef] [PubMed]
- 40. Iki, Y.; Ito, T.; Kudo, K.; Noda, M.; Kanehira, M.; Sueta, T.; Miyoshi, I.; Kagaya, Y.; Okada, Y.; Unno, M. Animal ethics and welfare education in wet-lab training can foster residents' ethical values toward life. *Exp. Anim.* **2017**, *66*, 313–320. [CrossRef]
- Redacción EFEverde. Suspenden la Actividad del Laboratorio Vivotecnia Tras ver Indicios de Maltrato Animal. 2021. Available online: https://n9.cl/zjdv6 (accessed on 11 May 2023).
- 42. Martínez-Sanchez, E. Implications of the Spanish Transparency Agreement on the use of animals in scientific research. *Lab. Anim.* **2017**, *46*, 33. [CrossRef]

- Pejman, N.; Kallas, Z.; Reig, L.; Velarde, A.; Moreno, M.; Magnani, D.; Protopapadaki, V.; Ribikauskas, V.; Ribikauskienė, D.; Dalmau, A. Should Animal Welfare be Included in Educational Programs? Attitudes of Secondary and University Students from Eight EU Countries. J. Appl. Anim. Welf. Sci. 2021, 26, 1–20. [CrossRef]
- 44. Aske, K.C.; Waugh, C.A. Expanding the 3R principles: More rigour and transparency in research using animals. *EMBO Rep.* 2017, 18, 1490–1492. [CrossRef]

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