1 2 3	Ethnopharmacology, Ethnomedicine, and Wildlife Conservation
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21	Abstract

Ethnopharmacological relevance: There are longstanding traditions of animal-derived products 22 23 being employed as medicines, and they continue to be important in many traditional cultural 24 healthcare practices. However, the populations of numerous so-used animals are known to be threatened with extirpation by such practices. Ethnopharmacological studies documenting these 25 26 animal-derived drugs are not only interesting from an anthropological standpoint, but they are also 27 relevant from a wildlife conservation perspective - especially since ethnopharmacologists are 28 intermediaries between indigenous and scientific communities, placing them at the forefront of being 29 able to ethically access information to address these issues.

*Methods:* Using the example of documenting culturally acceptable substitute materials for animal
 products (which ultimately also extends to flora), we explore the intersection of ethnopharmacology,
 biocultural resources, and wildlife conservation.

*Results:* Pharmacological efficacy and symbolism are factors influencing the utilization of traditional
 medicines. Achieving the integration of conservation aims with ethnopharmacology requires a
 nuanced understanding of both factors, along with fair adjudication when conservation and cultural

36 aims diverge. Ethnopharmacology is suitably placed for making conservation-orientated 37 recommendations - including investigating more sustainable substitutes for animal products in the 38 context of medical efficacy, and for engaging ethically with local communities to facilitate 39 information generation aimed at protecting the environment and their traditions.

40 Conclusion: We suggest an integrative approach to ethnopharmacological studies investigating 41 medicinal bioresource use. This approach is considerate of species' conservation profiles, the 42 substitutability and pharmacological efficacy of biocultural resources, indigenous and cultural rights, 43 and a collaborative ethos for stakeholder engagement.

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45 Keywords: Zootherapy, substitution, effectiveness, efficacy, animal conservation, health, medicine

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#### 47 **Graphical abstract**

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**1. Introduction** 



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67 The use of animal products driven by cultural practices, including traditional medicines, can lead to 68 unsustainable harvesting of species, and adversely impact wild populations and the welfare of animals 69 (e.g., Starr et al., 2010; Baker et al., 2013; Nijman and Nekaris, 2016). Animal products are used and 70 traded across a range of socio-economic and geographic contexts - from urban centres where 71 consumers are well-integrated in the global economy to informal rural markets and local subsistence 72 economies – and are integral to many complex cultural-spiritual traditions practiced by some of the 73 world's most vulnerable communities. Historically, ethnopharmacological field studies have aimed 74 to document such practices for purposes of scholarly inquiry, cultural preservation, and medical 75 insights. In a world where heterogeneous anthropological pressure is persistently exerted on wildlife 76 and other environmental resources, we present a series of considerations concerning the intersection 77 of ethnopharmacology and wildlife conservation - using the example of documenting substitute 78 materials for animal products. These considerations (akin to mindfulness) are not thorough; rather, 79 they serve as provocations for further debate and evaluation. By suggesting the integration of 80 conservation-considerate protocols into future study designs and publications, we hope to advance 81 the discourse on ways that ethnopharmacology can contribute meaningfully to biodiversity 82 conservation.

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### 84 **2. Methods and outline**

Amidst the heterogeneous human pressures impacting wildlife resources, we explored ideas concerning the intersection of ethnopharmacology and wildlife conservation. Proposed means for achieving wildlife conservation goals include, amongst other measures, supply side-interventions – encompassing substitute species, regulated trade, and wildlife farming (e.g., Bulte and Damania 2005; Phelps et al. 2014), enforced conservation laws leading to replacement and omittance of animal drugs from formulations (Yeshi et al., 2017), and databases proposing plant and mineral-based alternatives such as TAWAP (2024; https://tawap.org/).

92 Ethnopharmacologists can play important roles as intermediaries between indigenous and scientific 93 communities. They can also assist those conservation scientists who adopt a more wildlife population-94 centred research approach with collecting interdisciplinary data to: (i) understand the socio-cultural 95 use of species and their derivatives in a different context, (ii) inform population threat assessments 96 via the evaluation of the risks to species posed by some cultural practices, so that long-term 97 sustainability and availability for cultural use can be assessed and endorsed, (iii) report and test 98 culturally acceptable (and effective) substitutable materials, and (iv) report on instances of alien 99 species being adopted into pharmacopoeias (including as substitutes) to allow for a more nuanced 100 evaluation of the impacts of introduced species. These conservation-relevant considerations are not 101 exhaustive, and rather serve as prompts for engaging in further debate and evaluation.

We specifically considered here the role of ethnopharmacology in documenting substitutes and proposing more sustainable alternatives to animal products used in traditional medicines. We begin

104 by discussing the dynamics of traditional knowledge (section 3), followed by a discussion on 105 pharmacological efficacy (clinical setting), perceived effectiveness, and the symbolism (cultural 106 setting) of animals used in traditional medicines (section 4). In section 4 we also outline the use and 107 discontinuance of animal-derived materials based on their pharmacological efficacy in various parts 108 of the world and emphasise that use of animal-derived material is frequently driven by the symbolic 109 value and perceived effectiveness of the selected material. We thus describe two major (but not 110 exclusive) background considerations in the fair adjudication of potentially competing aims, i.e. that 111 of species to persist, and the rights of people to practice their traditions (section 5) – which we believe 112 are central to conservation-relevant (and socially just) practice. We suggest in section 6 that for 113 wildlife conservation elements to be integrated into ethnopharmacology research, careful 114 consideration should also be given to assessing the relative benefits and harms of the reported 115 practices - to wildlife and people (despite the inherent challenges in such assessments). In closing 116 (section 7), we emphasize the importance of considering equitable, collaborative, two-way 117 knowledge exchange between researchers and practitioners/users of traditional medicines - which 118 includes considering research instrument co-design, and information dissemination, with the study 119 participants, in accordance with ethical guidelines for conducting ethnobiological research 120 (Fernandez et al., 2003; International Society of Ethnobiology, 2006). By advocating for the explicit 121 integration of conservation-relevant considerations into future ethnopharmacology study designs and 122 publications, we aim to advance the discourse on how, within the ethnoscience disciplines, 123 ethnopharmacology can make substantive contributions to wildlife conservation initiatives and 124 interventions within the context of cultural practices - some of which may pose sustainability 125 challenges to important biological resources.

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### 127 **3.** The dynamics of traditional knowledge

128 In the transdisciplinary ethnosciences, indigenous and traditional knowledge are often considered 129 domains that *de facto* should be protected in their original context within communities (in situ) and 130 conserved outside of those communities through research and documentation (ex situ) for future 131 generations. However, these traditions of practice also lend themselves to periodic re-investigation 132 and review because culture and traditional knowledge are not static and unchanging (Cunningham 133 and Zondi, 1991). Instead, they are dynamic: i.e., capable of evolving and adapting to internal and 134 external pressures, including environmental changes, and can be transformed or abandoned (Bye et 135 al., 1995), and also deliberately invented (Hobsbawm and Ranger, 2012), leading to adaptations such 136 as the incorporation of new organisms and drugs into pharmacopoeias (e.g., non-native species) and 137 the lifting of cultural restrictions previously imposed on the use of taboo species (Cunningham and

138 Zondi, 1991). Strategies of enforced change have also been implemented through legislation. Culture 139 may also become formalized and institutionalized, thereby losing its dynamism while continuing its 140 expression as folklore. Historical evidence also indicates, however, that not all traditional knowledge 141 and customs are sustainable, and that not all traditions withstand the development of ethical 142 considerations, evolving moral standards, and/or critical evaluation of medical efficacy. 143 Consequently, some healthcare practices are abandoned following the discovery of more effective 144 alternatives, sustainable production methods and acquisition, or societal rethinking, sometimes 145 accompanying the implementation of new regulations protecting wildlife.

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### 147 4. Pharmacological efficacy and effectiveness of zootherapeutic remedies

Historically, the (more-or-less) systematic documentation of *materia medica* dates back, in some cultures, over 2000 years. These written sources include a taxonomically diverse array of crude drugs and preparations. Compendia, referred to as 'herbals' or 'herbal books', name the drugs they describe as 'herbal medicines' or 'herbal drugs', regardless of their biological or mineral origin. The custom of reporting animal drugs separately from botanical drugs is a more recent phenomenon (pers. obs., review of academic literature) and not constructive for understanding the consensus and use of *materia medica*.

155 Crude animal-derived 'herbal' drugs (i.e., zootherapeutic remedies prepared from the body-parts or 156 excreta of fauna) were an important part of these *materia medica* across Europe until the 18<sup>th</sup>/19<sup>th</sup> 157 centuries, and the teaching of 'pharmaceutical zoology' as a separate course was common in 158 European Pharmacy Schools during this period (see Plans y Pujol, 1870). The relatively uncritical 159 adoption and transmission of 'herbal' remedies facilitated the persistence of pharmacologically 160 ineffective drugs within *materia medica* over an extended period (MacKinney, 1946; Leonti, 2011).

161 Hence, just because herbal and traditional zootherapeutic medicines have been used for a long time 162 doesn't necessarily mean that their long-standing use serves as unequivocal evidence supporting the 163 medical efficacy and biomedical safety of treatments (Helmstädter and Staiger, 2014; EMA, 2024). Efficacy describes the capacity of an agent to produce an effect under standardized conditions, such 164 165 as in clinical trials, and effectiveness refers to the perceived therapeutic success in real-life practice 166 and within a cultural setting (Last et al., 2001, p. 57-58; Witt et al., 2013). Ethnopharmacological 167 research draws on anthropological data to critically evaluate knowledge regarding the therapeutic use 168 of traditional medicines by employing biological and chemical methods to test medical efficacy and

169 safety (Gertsch, 2009; Heinrich and Jäger, 2015; Weckerle, et al., 2018; Bruhn and Rivier, 2019).

As a consequence of the scientific revolution and advancements in rational thinking, a large number of European and Mediterranean Renaissance herbal drugs were subsequently systematically excluded

from pharmacopoeias and medicinal practices, starting in the 18<sup>th</sup> century (e.g., mumia powder from 172 173 ground mummies; Dannenfeldt, 1985). This exclusion was prompted by toxicological concerns and 174 a re-evaluation of their efficacy (Mann, 1984; Martins de Oliveira et al., 2019); crude animal-derived 175 drugs that had constituted a substantial portion of those materia medica were progressively excluded 176 from modern 'herbal' medicines. Matthioli's commentaries on Dioscorides' De Materia Medica 177 (Matthioli, 1967-1970), for example, included 76 chapters dealing with animal-derived drugs; 178 however, of these zootherapeutic drugs, only wool fat (lanolin), honey, and beeswax are currently 179 included in the European Pharmacopoeia 11.0 (2022). Additionally, in complementary and traditional 180 medicine (CAM), propolis, as well as milk and animal fat, are still in use. Furthermore, and to put 181 validated pharmacological efficacy into another context, only a limited number of traditional animal 182 drugs included in herbals have led to drug discoveries. For instance, the anticoagulant hirudin was 183 isolated from the saliva of the European medicinal leech (Hirudo medicinalis; Linnaeus, 1758) at the beginning of the 20<sup>th</sup> century. Leeches have been used for venesection and blood removal since the 184 2<sup>nd</sup> century AD as reported by the Greek scholar Galen; today, recombinant hirudin derivatives serve 185 186 as anticoagulant drugs (Nowak and Schrör, 2007). Chinese and Japanese medical traditions have 187 recommended the use of bear bile for biliary stone diseases for centuries (Hagey et al., 1993); the bile 188 of various vertebrate species, including humans, contains ursodeoxycholic acid, which can now be 189 produced semi-synthetically and used as a monotherapy for treating gallstone diseases (Hagey et al., 190 1993; Tonin and Arends, 2018). Another example of a potent animal-derived compound is the toxic 191 monoterpenoid cantharidin, found in Spanish fly (Lytta vesicatoria; Linnaeus, 1758), and other blister 192 beetles of the Meloidae family; cantharidin induces strong and long-lasting erections and has 193 therefore been used as an aphrodisiac or for treating erectile dysfunction (Pajovic et al., 2012).

194 Despite the exclusion of most crude animal-derived drugs from current herbalism, CAM, and official 195 pharmacopoeias in Europe and the US, animal derivatives remain an important source of allopathic 196 medicines (Zhu et al., 2011; Wagner et al., 2007; European Pharmacopoeia 11.0, 2022). These 197 derivatives are, however, obtained through biotechnological methods and via extraction from porcine, bovine, ovine, equine, mouse, and chicken products and organs (Wagner et al., 2007). A wide range 198 199 of animal-derived products are currently employed in the pharmaceutical and healthcare sectors, 200 including: anti-thrombotics, digestive supplements, lung surfactants, haemostatic agents, vaccines, 201 anti-diarrheal agents, steroids, pituitary and gonadal hormones, plasma volume expanders, immune 202 supplements, immunomodifiers, anti-neoplastics, anti-migraine, anti-hypertensive and hematopoietic 203 agents, ophthalmic medications, anti-rheumatoid agents, anti-venoms, and anaesthetics (Zhu et al., 204 2011; Wagner et al., 2007; Queensland Health, 2020). Furthermore, derivatives from marine invertebrates and arthropods also contribute to drug discovery by serving as valuable sources of
highly bioactive secondary metabolites (Kiyota, 2021; Dossey, 2010; Seabrooks and Hu, 2017).

207 But while there has been a significant reduction in the number of animal-derived drugs in European 208 and American pharmacopoeias, there has been little such significant reduction in many traditional 209 medicine practices and the extensive zootherapeutic use of animal products remains important to 210 many traditions of socio-cultural medicine across the world – especially in countries where traditional 211 medicine has a strong presence or in communities which depend less on, or who have less access to, 212 western-based primary healthcare. For example, Moorhouse et al. (2022a) reported over 2300 species 213 of wild fauna used in traditional Chinese medicine. Additionally, a survey conducted at a traditional 214 medicine market in Johannesburg, South Africa, identified 147 traded vertebrate species (Whiting et 215 al., 2011). A review based on market surveys conducted in 20 Brazilian cities documented animal 216 medicines deriving from 131 species (however, the actual number of taxa used throughout Brazil is 217 likely much higher) (Ferreira et al., 2013). Closer to Europe, Bellakhdar's (2020) book on the 218 traditional pharmacopeia of Morocco details 227 animal taxa products across 89 chapters.

219 In contrast to cases where animal-derived medicines have demonstrable medical properties, a 220 considerable proportion of cultural zootherapeutic preparations are pharmacologically inactive 221 (MacKinney, 1946; Still, 2003) and lack medical efficacy, yet are considered by users to be highly 222 effective medicines for treating the recognised disorders. Animal derivatives constitute integral 223 components of complex socio-cultural practices that rely on the symbolic, and often non-224 consumptive, spiritual and characteristic properties of the animals involved where bodily traits and 225 functions signify utility (Williams and Whiting, 2016). Explanations for the use of crude animal 226 products have included beliefs that the biological and spiritual characteristics and qualities of the 227 animals are transmitted to patients, following the principles of the Doctrine of Signatures (De 228 Conconi and Moreno, 1988; Lev, 2002; Fresquet Febrer, 2001). For example, in southern African 229 traditional and spiritual practices, lion (*Panthera leo*; Linnaeus, 1758) parts are used symbolically to 230 confer bravery, strength, and power to the user, and are thus closely associated with traditional royalty 231 (Simelane and Kerley, 1998; Coals et al., 2022). This association may be manifested through the 232 predominantly non-consumptive use of lion claws, teeth, bones, and skin - often as part of tribal 233 regalia (Coals et al., 2022; Williams et al., in review). Furthermore, animals have been noted to be 234 used for treating disorders that are difficult to define in Western medical concepts and/or are based 235 on healers' interpretations of causation (some of which are also social rather than medical disorders) 236 (Simelane and Kerley, 1998; Simelane, 2011). Thus, ethnopharmacologists do not typically consider 237 testing the therapeutic efficacy claims associated with animal-derived traditional medicines, such as 238 of the efficacy of charcoaled hedgehogs (Budjaj et al., 2021), tiger's breastmilk, or the whole dried

body of the slow loris (Chassagne et al., 2016), for practical, ethical, and above all, epistemologicalreasons.

241 We therefore contend that the symbolic and spiritual use of animal products in traditional therapeutic 242 practices is likely to outweigh the active-ingredient-driven (modern 'Western') conceptions of 243 medicine, and we thus caution against simplistic interpretations of medical efficacy and effectiveness. 244 This raises questions about the degree to which knowledge of the traditional uses of animal products 245 contributes to drug discovery, while acknowledging the need for case-by-case differentiation. While 246 the documentation of animal-derived medicines has the potential to contribute to the preservation of 247 cultural diversity and the development and implementation of protection measures, much of this 248 knowledge is unlikely to meaningfully advance medicinal discoveries for future generations. 249 Nevertheless, symbolism and perceived effectiveness are key drivers in the enduring use of many 250 cultural-spiritual medicines.

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# 252 5. Integrating conservation concerns into ethnopharmacology requires evaluation of competing 253 needs

254 While having baseline data about the use and trade of traditional medicines can help inform 255 conservation strategies, the practical implementation (i.e., translational measures) of these strategies 256 is often uncertain and beyond the direct control of academic researchers. In light of this, we therefore 257 suggest that, where possible, academic researchers consider making more direct and proactive 258 contributions to conservation endeavours. A short discussion between Nijman and Nekaris (2016) 259 and Chassagne (2017) about reporting animal drugs used in traditional medicine, for example, 260 published in this journal, may not have received the attention it deserved. Nijman and Nekaris (2016) 261 emphasized that the conservation status of the respective animal species should be reported, 262 regardless of whether the animal needs to be killed to obtain the part used as a drug or not. 263 Furthermore, they stressed that the legality of collecting and trading the respective products should 264 also be stated (it should be noted that these statements would be aimed at readers of the journal reports 265 and not intended as parochial commentary to indigenous users). Nijman and Nekaris (2016) further 266 reminded researchers to "provide context about the ethical and legal implications" when reporting 267 on the use of protected and endangered wildlife in ethnopharmacological surveys (see also 268 Chassagne, 2017). In response to Nijman and Nekaris (2016), Chassagne (2017) pointed out that, 269 according to Article 24 of The United Nations Declaration on the Rights of Indigenous Peoples 270 (UNDRIP), "indigenous peoples have the right to their traditional medicines and to maintain their 271 health practices, including the conservation of their vital medicinal plants, animals and minerals". 272 Moreover, contributing to the conservation and sustainable use of biodiversity is an integral aspect of The Journal of Ethnopharmacology (see: <u>https://www.sciencedirect.com/journal/journal-of-</u> ethnopharmacology/about/aims-and-scope) and encapsulated in a founding objective that investigations of agents used in traditional medicines should not exploit them (Bruhn and Rivier, 2019).

277 Whilst there may be a generally held desire to honour traditions and cultural practices that may 278 involve the use of animal products, when such practices conflict with conservation aims, particularly 279 the potential persistence of a species, an evaluation of those competing needs is required (Coals et 280 al., 2019). Such evaluations necessitate an acknowledgement of the potential influences and pitfalls 281 of cultural relativism, whereby a potentially damaging activity may be overlooked because of its perceived cultural significance (Dickman et al., 2015). We posit that a de facto position of the 282 283 assumed right to preserve all cultural practices, as presented in many ethnobiological publications, 284 may serve to perpetuate potentially harmful practices, which can ultimately lead to the extinction of 285 fauna and traditions. Therefore, enhancing the integration of ethnographic studies with conservation 286 requires evaluating conflicting interests and providing an explicit and objective rationale for 287 preserving practices involving at-risk wildlife (Macdonald et al., 2021), coupled with recognizing 288 that cultural practices and conservation aims will not always be compatible but that they should be 289 resolved in a fair and just manner (Vucetich et al., 2018).

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### 291 6. Substitution is not simple but should be considered

Academic research on the utilisation of traditional medicine should not confine itself only to documenting therapeutic uses for wildlife, as use-based demand has negative consequences for many species. Hence, the cultural and ecological aspects of these practices should also be explored (Alves and Rosa, 2007).

296 The potential substitution of wild-sourced preparations with those from domestic animals or botanical 297 materials has become a topic of considerable interest in some primarily conservation-focused 298 publications. However, the prevalent use of preference-based surveys and techniques, upon which 299 many findings are based, tends to overlook complexities in real-world markets and dichotomies 300 between what people will say and what they do (Hinsley and 't Sas-Rolfes, 2020). While individuals 301 may express preferences for potentially effective substitutes in experimental and hypothetical 302 scenarios (where effectiveness is not synonymous with efficacy), the conditions for acceptance of 303 such substitutes is uncertain, primarily due to the significant cultural connotations associated with the 304 use of specific materials (Ferreira et al., 2016). For example, although several, allegedly effective, 305 symbolic alternatives ostensibly exist for lion products in southern African spiritual practices, aspects 306 of the lion's role in spiritual healing have been found to be largely immutable and, furthermore, that

ancestral spirits (considered divine messengers between the healers and a higher being) may, during dreams and rituals, be influential in the suggested prescription of some effective alternatives (Coals et al., 2022). Moreover, while botanical substitutions are increasingly widely touted as substitutes by researchers with different world views to the consumers, care must be exercised to ensure that they (and faunal alternatives) are also not sourced from threatened species. Hence, we caution against naïve assumptions that botanical alternatives will automatically be the more acceptable, conservationfriendly, options.

314 In instances where substitution is a matter of efficacy (e.g., Moorhouse et al., 2022b), 315 pharmacological studies are well-placed to aid in the identification of effective and sustainable 316 substitutes. However, pharmacological efficacy has rarely been established for animal-derived 317 traditional medicines (see above) and clinical trials for this category are generally lacking (Simelane 318 and Kerley, 1998). Conversely, efficacy and safety have been established for numerous botanical 319 drugs, while for others, their efficacy is considered plausible (e.g., EMA, 2024). Thus, available 320 pharmacological and clinical literature can be used to assess evidence of substitute efficacy on a case-321 by-case basis.

322 We emphasize that animal-based ethnopharmacological field studies should consider documenting substitutes (sustainable, effective, non-threatened, or otherwise) (Chassagne, 2017). This 323 324 documentation could be an outcome of questions on substitutes specifically incorporated into the 325 research instruments from the outset, or from information provided unprompted by stakeholders 326 during the course of the research process. Such information is held by traditional healers, consumers, 327 and sellers. The intention is not to question the rights of indigenous peoples to choose their medicines. 328 Rather, we advocate for the documentation of culturally acceptable substitutes (where they exist, and 329 the traditional knowledge has been obtained through informed and consensual ethical research) as 330 part of collaborative systems with communities to address conservation challenges through mutual 331 feedback (for an example concerning the adoption of faux leopard skins by followers of the Shembe 332 Church in South Africa see: Naude et al. 2020). Thus, this approach is not about overriding indigenous peoples' rights to maintain their traditional health-care practises, but rather to actively engage with 333 local communities to facilitate taking steps towards information generation aimed at protecting the 334 335 environment and their traditions.

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### **337 7. Considering the future of ethnopharmacological reporting**

In addition to the ongoing global changes that exert pressure on biodiversity, some traditional medicine practices contribute to the decline and extirpation of some animal populations – thereby posing further threats to the persistence of species. Although users may assert the effectiveness of 341 animal-derived medicines, the strictly therapeutic efficacy of most of these medicines has not been 342 tested and remains questionable, with use often driven by complex, culture-specific, traditions of 343 practice, ancestral guidance, and symbolic connotations. Consequently, we suggest that field 344 researchers adopt a collaborative approach to explore the possibility of establishing consensus 345 regarding culturally accepted alternative materia medica (of whatever origin) especially those that 346 can be sourced from non-threatened species that can be harvested more sustainably. A first step would 347 be to investigate the nexus of traditional uses of biological and inorganic substances used by specific 348 communities so that possible convergent uses and consensus can be pinpointed, and substitutability 349 can be assessed in collaboration with healers, sellers, and consumers. Feedback on more effective 350 substitutes and alternatives are more likely to be accepted when proposed in a culturally respectful 351 and inclusive manner, particularly by considering the culture-specific symbolism of the species 352 intended for replacement. While indigenous cultures, rural and urban populations have the right to 353 maintain their traditional health practises, we contend, however, that they have also the right to be 354 kept apprised of the scientific evaluations of their customs and of the environmental impacts of their 355 practices. Thus, within the domain of 'conservation-conscious ethnopharmacology', we suggest an 356 integrative approach to ethnopharmacological studies investigating medicinal bioresource use (Fig. 357 1). This approach is considerate of species' conservation profiles, the substitutability and 358 pharmacological efficacy of biocultural resources, respect for indigenous and cultural rights, and a 359 collaborative ethos for stakeholder engagement.

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Figure 1. Considerations for integrating wildlife conservation-conscious factors into
 ethnopharmacology research, using the example of potential substitutions for animal products.

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