- 1 Title: Diversity and evidence gaps among potential win-win solutions for conservation and
- 2 human infectious disease control
- 3 **Running Title:** Win–wins for conservation and infectious disease control
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### 60 Summary:

As sustainable development practitioners have worked to "ensure healthy lives and promote well-61 62 being for all" and "conserve life on land and below water," what progress has been made with 63 win-win interventions that reduce human infectious disease burdens while advancing conservation 64 goals? Using a systematic literature review, we identified 46 such proposed solutions, which we then investigated individually using targeted literature reviews. The proposed solutions addressed 65 diverse conservation threats and human infectious diseases, and thus the proposed interventions 66 67 varied in scale, costs, and impacts. Some potential solutions had medium- to high-quality evidence for prior success in achieving their proposed impacts in one or both sectors. However, there were 68 69 notable evidence gaps within and among solutions, highlighting opportunities for further research 70 and adaptive implementation. Stakeholders seeking win-win interventions can explore this review and an online database to find and tailor a relevant solution or to brainstorm new solutions entirely.

71 72

Keywords: biodiversity, disease ecology, ecosystem health, epidemiology, intervention, One
 Health, nature-based solutions, public health, restoration

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## 76 Main Text:

77 Introduction

78 Ecosystem degradation can exacerbate infectious diseases that have long plagued humankind or

cause novel pathogens to spill over from animals to humans.<sup>1–6</sup> By targeting connections between human infectious disease and the natural world, interventions might both "ensure healthy lives and

human infectious disease and the natural world, interventions might both "ensure healthy lives and
promote well-being for all" and "conserve life on land and below water"—two Sustainable

2 Development Goals (SDGs).<sup>7-17</sup> For example, putting tick collars on free-ranging dogs might

reduce transmission of ticks and tick-borne disease from dogs to people and wildlife.<sup>18</sup> Indeed,

sustainable development practitioners around the world are urgently seeking safe and effective

85 cross-sector interventions that might prevent the next pandemic.<sup>19</sup>

86

Of course, no single win–win intervention will work in all contexts or solve all problems within complex socio-ecological systems.<sup>20</sup> Interventions that improve some outcomes for human health and ecosystems might even cause collateral impacts in other sectors, creating complex trade-offs among SDGs.<sup>16,21,22</sup> Tasked with choosing an optimal intervention for any given problem and socio-ecological context, practitioners need to know which intervention options exist and how to compare them. Or, in the event that no existing intervention is suitable, practitioners will need to know how to identify and evaluate new intervention options.

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95 Unfortunately, the information needed to identify, implement, and evaluate win-win interventions 96 that prevent or control human infectious diseases tends to be limited, inconsistent, and/or 97 unconsolidated.<sup>23</sup> For instance, among conservation intervention studies that reported human well-98 being benefits, fewer than 2% considered health-specific outcomes, and only a subset of those considered emerging or endemic infectious diseases.<sup>24,25</sup> Furthermore, existing studies are 99 scattered across siloed disciplines that use different research methodologies, measure different 100 101 outcomes, and publish in different journals. Navigating this dispersed evidence landscape would 102 be prohibitively time-consuming for practitioners interested in implementing win-win 103 interventions.

105 To facilitate timely, evidence-based decision-making, we review existing evidence regarding win-106 win solutions that aim to simultaneously reduce human infectious disease burdens and advance 107 conservation goals. Because there are growing policy initiatives for enacting "nature-based 108 solutions",<sup>8</sup> we use the terms "intervention" and "solution" interchangeably throughout. First, we 109 provide a "menu" of 46 such proposed solutions ranging from local to international scales and 110 their individual information summaries. Stakeholders can explore these examples to find and tailor 111 potential solutions to meet their needs, or to brainstorm new solutions entirely. We then synthesize 112 information across the 46 solutions to describe some general criteria that stakeholders can use to 113 identify and evaluate potentially viable solutions that achieve their specific goals within their 114 resource constraints. Finally, we highlight evidence gaps within and among solutions that could 115 be important targets for future research and implementation.

116

### 117 Search Strategy and Selection Criteria:

118 To find and synthesize evidence among proposed solutions, we used a subject-wide evidence synthesis, a two-phase approach for identifying and assessing a broad suite of interventions 119 supported by heterogeneous evidence (Fig. 1).<sup>26,27</sup> In the first phase, we performed a systematic 120 literature review of peer-reviewed papers and book chapters, which we used to identify solutions 121 that have been proposed to simultaneously reduce human infectious disease burdens and advance 122 123 conservation goals (see below; appendix p 2-4). In the second phase, we performed targeted rapid 124 reviews<sup>28</sup> of the peer-reviewed and gray literature for each proposed solution, iteratively revising 125 evidence summaries for each (see below; appendix p 4-5). Finally, we used these evidence 126 summaries to categorize information for each solution, making it easier to synthesize and compare 127 among solutions.

128

129 To create a list of proposed win-win solutions (Phase 1), we systematically reviewed publications in Thomson Reuters Web of Science<sup>TM</sup> (http://thomsonreuters.com) and NCBI Pubmed® 130 (https://pubmed.ncbi.nlm.nih.gov) on 14 March 2018 (N=12,270 papers), including records 131 132 published any time prior. We performed the search using 167 English search terms regarding 133 conservation, ecology, infectious disease, and human populations (adapted from McKinnon et 134 al.<sup>25</sup>; see appendix p 2-4 and 14 and Fig. S1 for PRISMA diagram). We identified 617 papers containing hypothesized or measured outcomes for both conservation and human infectious 135 136 diseases—excluding papers that lacked proposed outcomes for one or both sectors—by using a 137 combination of researcher classification and machine learning to sort records by relevance<sup>29</sup>. 138 During subsequent full-text analysis, we removed any records that did not suggest at least one 139 proposed win-win solution (e.g., papers about trade-offs where environmental degradation 140 improves health). We then used full-text analysis of the final list of 383 records to group records 141 pertaining to the same win-win solutions into collective case studies (appendix p 4). This resulted 142 in a list of 46 unique proposed solutions (Fig. 2).

143

Each solution was then individually reviewed by one or two investigators (Phase 2: targeted rapid reviews), where keyword searches were used to find additional peer-reviewed publications on Web of Science<sup>TM</sup> and online searches were used to find gray literature. These rapid reviews were not systematic because they did not examine all published literature—a task that would not be possible for 46 interventions. Instead, investigators specifically sought publications with information relevant to 20 information categories, determined *a priori*, and summarized all information in a

150 standard format (appendix p 4-5). A single lead investigator reviewed all collective case study

151 summaries to ensure consistency, and then each summary was reviewed by an external expert

152 (appendix p 5). Based on feedback from external experts, investigators iteratively searched for

- 153 more information and revised the collective case study summary until the investigator and lead
- 154 investigator deemed the review complete. The 46 summaries are available in an open access
- 155 database: https://ecohealthsolutions.stanford.edu/research/win-win-solutions-people-and-nature
- 156

157 After finishing the collective case study summary, each investigator used a list of qualitative 158 variables defined a priori to categorize information in a consistent way that could be compared 159 across case studies. These variables included geographical location, conservation threat, infectious 160 disease threat, mechanism or "lever" type, evidence for conservation and human infectious disease 161 outcomes, and 11 criteria that we identified as indicative of viable solutions: Harmless, Contained, Consistent, Feasible, Acceptable, Impactful, Effective, Affordable, Scalable, Sustainable, and 162 163 Cost-Effective (see definitions in appendix p 7-8). The investigators' designations were confirmed 164 by the lead investigator and 1-2 other investigators to ensure consistency. Any discrepancies between how different people categorized information was discussed until consensus was reached. 165

166 After information from all proposed solutions had been categorized, we synthesized information

167 across the 46 solutions to describe their diversity and evidence gaps (Figs. 2-6).

168

169 *Potential solutions were widespread and diverse:* 

The 46 potential solutions addressed diverse threats, collectively covering all continents (except 170 Antarctica), most major pathogen groups (except fungi), and most conservation threat classes 171

172 defined by the International Union for the Conservation of Nature (except "geological events" and

"other"; Fig. 4).<sup>30</sup> Most solutions addressed multiple threats for health and conservation (e.g., 173 multiple pathogen species, multiple IUCN threat classes). The 46 potential solutions also covered 174

175 many intervention types and targets, ranging from vaccinating vampire bats against rabies in Peru<sup>31</sup>

to establishing sustainable harvesting programs for medicinal plant species in Tanzania.<sup>32</sup> Potential 176

177 solutions were diverse because the problems that they addressed were diverse.

178

179 Most solutions addressed pathogens with environmentally mediated transmission, such as vector-180 borne diseases and zoonotic diseases transmitted from animals to people (Fig. 4A), mirroring the

strong focus on these diseases in the One Health, Planetary Health, and EcoHealth fields.<sup>9,33,34</sup> For 181

182 example, the World Health Organization (WHO) and other international organizations support 183 expanding training for integrated pest and vector management globally, because these

184 management techniques might reduce total pesticide use while controlling both crop pests and 185 disease vectors such as mosquitoes. Environmentally mediated diseases like these have probably been the easiest entry points for cross-sector solutions due to their obvious underlying links 186 187 between human health and ecosystems.

188

189 Only a few potential solutions addressed diseases without environmentally mediated transmission, 190 like HIV/AIDS and pneumonia. For example, people in poor health due to HIV/AIDS or other 191 diseases are more likely to use easier and more destructive fishing practices in communities near 192 Lake Victoria, so HIV/AIDS treatment and prevention programs might support both human health

193 and aquatic conservation.<sup>35</sup> Focusing on these understudied links between ecosystems and directly

194 transmitted and chronic human diseases might yield additional solutions for advancing 195 conservation and health.

197 In addition to addressing diverse health threats, the potential solutions also addressed all IUCN 198 conservation threats related to anthropogenic activities (Fig. 4C). Solutions related to land use

199 change—agriculture & aquaculture (n=29) and urbanization & development (n=22)—were most

200 common, probably because land use change is a leading driver of biodiversity declines<sup>36</sup> and

- disease spillover.<sup>6,33,37,38</sup> In contrast, only a few solutions addressed transportation corridors,
   energy and mining, and climate change, and never as the primary threat (Fig. 4C).
- 203

We did not include a collective case study where climate change was the primary conservation threat and global emissions reduction was the solution (appendix p 3-4), because the many health and conservation outcomes that could be achieved by global emissions reduction<sup>9,10</sup> have yet to be measured. Climate change is expected to become a more urgent threat over time,<sup>7</sup> so there is a clear need for actionable, targeted solutions related to climate change.

209

210 The conservation and health threats targeted by the potential solutions spanned all geographical scales, from sub-national or national extents to regions (multiple countries within a continent) to 211 212 multiple continents (Fig. 4A). For example, vaccinating prairie dogs to reduce plague risk for 213 endangered black-footed ferrets and humans applies sub-nationally to the Western United States,<sup>39</sup> 214 whereas forest conservation to reduce human malaria might be relevant to multiple countries in Latin America, Africa, and Asia.<sup>40,41</sup> The potential solutions included targets in low- and middle-215 216 income countries, where there are high burdens of environmentally mediated human infectious 217 diseases, and targets in high-income countries, where infectious disease burdens are lower and 218 research effort is higher (Fig. 4A). Ultimately, one or more potential solutions probably exist for 219 all countries, but the set of relevant solutions that apply to any given country could be expanded 220 by future efforts to scale or translate existing solutions to new locations.

221

222 There were 27 potential solutions that involved implementing classic conservation interventions that have health benefits ("conservation levers for health"; Fig. 3), which are sometimes called "nature-based solutions" or "ecological levers for health".<sup>8,42,43</sup> One particularly common 223 224 225 conservation intervention type was species management, such as controlling or eradicating 226 invasive honeysuckle to reduce negative impacts on native vertebrates and reduce vector populations associated with honeysuckle,<sup>44</sup> and reintroducing native prawns extirpated by dams to 227 help control the snails that transmit human schistosomiasis.<sup>45</sup> Another common conservation 228 intervention type was land or water management or protection, such as conserving or restoring 229 230 wetlands to restore biodiversity while also reducing waterborne diarrheal diseases.<sup>46</sup> Together, 231 these 27 potential solutions are the most comprehensive list to date of conservation solutions that 232 might specifically reduce human infectious diseases, an important subsector within the global focus on conservation solutions that improve general human well-being.<sup>8,9,33,42</sup> 233

234

235 Six potential solutions involved classic public health interventions that have conservation benefits 236 ("health levers for conservation"; Fig. 3). For example, health system strengthening or family planning and reproductive health programs-including Population, Health, and Environment 237 (PHE) programs in many countries<sup>47</sup>—have reduced illegal logging and deforestation,<sup>48,49</sup> 238 improved coral and mangrove conditions in marine environments,<sup>50</sup> and improved community 239 participation in or approval of conservation initiatives.<sup>51</sup> There were also several solutions that 240 used insect vector control to reduce vector-borne disease risk for both people and wildlife.<sup>52,53</sup> 241 242 These health interventions were often supported by limited evidence, either because there were

not enough resources dedicated to monitoring and evaluation, or because interventions
implemented by the health sector did not quantify ecosystem or conservation outcomes. Therefore,
we expect that more health interventions that advance conservation goals might be imagined
through increased collaboration between conservation and health organizations.

247

248 Finally, 13 potential solutions acted through interventions that were neither specific to public 249 health nor conservation, but which affected both sectors ("levers for health and conservation"; Fig. 3). Many of these were policies regarding the food-energy-water nexus,<sup>54</sup> such as regulating 250 protozoan pollution,<sup>55</sup> reducing antibiotic use in aquaculture,<sup>56</sup> reducing nutrient pollution and 251 eutrophication associated with agriculture,<sup>57</sup> and implementing ballast water treatment protocols 252 to prevent invasive pathogens and wildlife from moving among ports.<sup>58</sup> There were also outreach, 253 254 education, or livelihood interventions, such as teaching people how to live safely with bats that 255 might be virus reservoirs;<sup>59</sup> protecting tree sap collection pots from bat contamination using bamboo skirts;<sup>60,61</sup> and replacing wood-burning stoves with cleaner cookstoves to reduce 256 deforestation and smoke-related pneumonia.<sup>62</sup> However, livelihood-focused interventions were 257 258 relatively rare, so future efforts might discover more interventions that primarily target poverty 259 and inequalities (SDGs 1 and 10) and that have downstream benefits for health and conservation 260 (SDGs 1, 14, and 15).

261

The diversity among potential solutions is promising, because "there is no one-size-fits-all 262 approach for One Health implementation."<sup>33</sup> The 46 examples described here cover many context-263 264 specific health and conservation threats, so stakeholders might be able to adapt one of these to meet their needs. Where none of the 46 potential solutions described here are relevant, stakeholders 265 266 could design new solutions to meet their specific goals within their resource constraints. To 267 determine whether any given solution will be viable for a given context, stakeholders can evaluate 268 the 11 viability criteria described below: Harmless, Contained, Consistent, Feasible, Acceptable, 269 Impactful, Effective, Affordable, Scalable, Sustainable, and Cost-Effective (Fig. 5, also see 270 appendix p 7-8). 271

272 Identifying and minimizing trade-offs:

273 To evaluate potential trade-offs caused by a given intervention, we suggest considering three 274 viability criteria (Figs. 2, 5): Harmless solutions are not expected to harm non-target aspects of 275 human well-being for some people while attempting to help other people; *Contained* solutions are 276 not expected to have negative, collateral impacts on non-human targets, or else potential collateral 277 impacts could be avoided or fully mitigated; and Consistent solutions are expected to have only 278 positive outcomes for their intended conservation and human infectious disease control targets in 279 predictable contexts (i.e., no known negative outcomes). Two investigators used these definitions 280 and available evidence to determine whether each potential solution met, failed, or was data-281 deficient for the three criteria (also see appendix p 6-7). Data limitations often made it difficult to 282 decide whether a solution involved substantial trade-offs, but existing evidence demonstrated that 283 19 solutions could be *Harmless*, *Contained*, and *Consistent* under some contexts (Figs. 1 and 5).

284

285 Context-dependency was common among the 46 potential solutions. For example, introducing

286 invasive predators that consume larval mosquitoes to control malaria has often negatively

- 287 impacted ecosystems (e.g., violating the *Contained* criterion),<sup>63</sup> but in at least some contexts
- 288 (e.g., native predators), mosquito predators are expected to have only positive outcomes for

- 289 ecosystems (meeting the *Contained* criterion). However, for other solutions, we could not
- 290 identify a mediating context. For example, correlational studies suggest that forest cover and host
- biodiversity impact competent host abundance in ways that can reduce human Lyme disease risk
- in North America (the "dilution effect").<sup>64</sup> Yet in this complex system, forest cover or host
- biodiversity have also been associated with unexpected amplification of Lyme disease risk<sup>65</sup>, and
- thus it is unclear how forest conservation or restoration interventions would impact human health
- 295 (i.e., this solution violates the *Consistent* criterion). Future research or innovation might identify 296 specific, predictable scales and circumstances where this solution does no harm, but at present, it
- risks causing unpredictable harm to some people in some contexts.
- 298

299 Like most solutions, those that address pathogen spillover from wildlife to humans had data gaps 300 regarding trade-offs. For example, wildlife trade is a conservation threat and a pathway for spillover from wildlife.<sup>66-69</sup> However, in several African countries, past bans on all wildlife 301 302 hunting and consumption sometimes created food insecurity, illegal markets, and distrust in health authorities.<sup>21,70-72</sup> Bans developed in response to the recent COVID-19 pandemic might be 303 similarly problematic.<sup>73,74</sup> This evidence demonstrates that wildlife trade bans can cause harm 304 305 when they affect subsistence hunting and consumption. But wildlife trade bans or restrictions 306 might be safe and feasible in other specific contexts. For example, many existing national and 307 international wildlife pet trade restrictions aim to conserve wildlife (e.g., the Convention on 308 International Trade in Endangered Species; https://cites.org), and there is increasing (but not 309 universal) public support for bans or restrictions on "luxury" commercial wildlife trade in Asia 310 due to the recent coronavirus outbreaks.<sup>75</sup> If successful, restrictions and bans that target the multibillion-dollar commercial wildlife trade could prevent multi-trillion-dollar pandemics.<sup>16,76</sup> 311 312 However, it is still unclear whether and when these policies can be successful, including whether 313 they will favor illegal markets or erode support for conservation. Efforts to identify contexts where 314 negative impacts are mediated, conservation is advanced, and spillover risks are reduced are 315 urgently needed.

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### 317 Achieving Socially Acceptable and Feasible solutions:

Two criteria can be used to determine whether solutions are immediately achievable (Fig. 5): 318 319 Feasible solutions could be successfully implemented now, given existing technology and 320 sufficient resources, and *Socially Acceptable* solutions are supported by the stakeholders who are 321 affected by the intervention or could be made acceptable to the stakeholders. For example, the 322 livestock medication diclofenac was implicated in dramatic vulture population declines in India, 323 which might have allowed free-ranging dog populations and human rabies risk to increase.<sup>77</sup> 324 Therefore, diclofenac was banned in India and surrounding nations, an intervention that was 325 achievable because acceptable, alternative veterinary drugs that could replace diclofenac and that 326 were not toxic to vultures already existed.<sup>77</sup> As this example illustrates, some of the 46 potential 327 solutions have already been successfully implemented on national or multinational scales.

328

329 However, most potential solutions were data deficient for feasibility or social acceptability. For

example, two potential solutions involve broadly "rewilding" North America with top carnivores

to control infectious diseases in wild herbivores and possibly humans.<sup>78</sup> These potential solutions

- face opposition from some stakeholders (e.g., ranchers, hunters), and it is unclear whether
- rewilding can be made socially acceptable in places where the intervention would most impact
- 334 disease transmission (or whether rewilding would lead to net reductions in disease risk). Indeed,

335 solutions that involved changing peoples' lifestyles and cultures were often data deficient for 336 *Acceptability*. This highlights a clear need for future human-dimensions and implementation 337 research to evaluate cross-sector solution viability, as outlined by the Organisation for Economic

- 338 Cooperation and Development (OECD).<sup>79</sup>
- 339

#### 340 Impactful and Effective solutions achieve stakeholders' goals:

341 Whether a given solution meets stakeholders' goals can be evaluated using two criteria: Impactful 342 solutions have the potential to meet stakeholders' quantitative goals (i.e., effect magnitude, clinical 343 relevance), and *Effective* solutions can successfully achieve the desired outcomes. For example, 344 building nest boxes to increase local predatory bird populations is proposed to control the rodent species that are reservoirs for hantaviruses.<sup>80</sup> This is an ecological lever for health,<sup>42</sup> but there is 345 346 no evidence that nesting sites are limited in this example, so the intervention may only re-distribute 347 non-threatened wildlife populations in ways that benefit humans, creating little value for 348 stakeholders with strong conservation priorities (i.e., this solution might not be Impactful for 349 conservation). There is also limited evidence that this solution can successfully reduce human 350 disease burdens (i.e., this solution might not be *Effective* for human health). We did not quantify 351 how Impactful each potential solution was, because there was no common system available to rank 352 impacts given the diverse methods and metrics used across the relevant literature. However, we 353 did qualitatively assess how *Effective* each solution was, as evidenced by prior success in achieving 354 proposed goals (Fig. 6, appendix p 5-7).

355

356 We categorized existing evidence quality for each potential solution using a modified Bridge Collaborative rubric<sup>81</sup> with three categories: evidence types and diversity, evidence consistency, 357 358 and evidence applicability (appendix p 5-7 and 15, Table S1). We then combined these three 359 categories into one composite score for overall evidence quality for health outcomes and one score 360 for conservation outcomes (Figs. 2 and 6): "no evidence" indicated that cases were supported only by hypotheses and anecdotes; "low evidence quality" indicated that there were limited supporting 361 362 studies with moderate to major evidence gaps, unexplained inconsistency, or limited applicability; 363 "medium evidence quality" indicated that there were several lines of evidence that were mostly 364 consistent and applicable, where inconsistency could be explained; and "high evidence quality" 365 indicated diverse evidence types, usually including an intervention study, that yielded consistent 366 and applicable results and left little to no uncertainty regarding the outcome. The resulting 367 composite evidence quality scores highlight which solutions have had demonstrable success for 368 health and conservation outcomes and which still have evidence gaps.

369

370 There were seven solutions that already had medium to high evidence quality for both conservation 371 and human health success and were thus *Effective* (Fig. 6). For example, vaccinating dogs and wild 372 carnivores to reduce rabies transmission among dogs, wildlife, and people was supported by high evidence quality for both outcomes, including successful intervention programs.<sup>82,83</sup> Most 373 solutions had higher evidence quality regarding *Effectiveness* for conservation than for health. For 374 example, it is well established that controlling invasive rats, brushtail possums, and cats can 375 conserve endemic species,<sup>84-86</sup> especially on islands. There are also studies linking invasive 376 species control and human infectious disease burdens,<sup>87,88</sup> but the evidence types and diversity are 377 more limited. Similarly, though forest restoration and conservation have well-established benefits 378 to ecosystem structure and function,<sup>89,90</sup> and several correlational studies link upstream forest 379 cover to reduced childhood diarrhea risk downstream,<sup>91</sup> there are no intervention studies linking 380

forests to childhood diarrhea. Though evidence for these solutions could still be improved, these examples show that *Effective* cross-sector solutions do exist.

383

384 There were 17 potential solutions that had low overall evidence quality due to low evidence 385 diversity, low evidence applicability, and/or evidence inconsistency that was difficult to explain 386 for one or both outcomes. For example, a few observational studies (limited evidence diversity) 387 quantified high leopard predation rates on free-ranging domestic dogs, an important disease reservoir for rabies near Mumbai.<sup>92</sup> Leopard conservation might reduce dog rabies and thus human 388 rabies risk, but predation or culling of dogs could also counterintuitively increase rabies in dog 389 reservoir populations (potentially not Consistent).<sup>93</sup> In another example, regulating drawdown 390 rates for water reservoirs created by dams might restore aquatic communities and reduce larval 391 mosquito survival and thus human malaria risk near dams.<sup>94</sup> However, the existing evidence did 392 393 not come from countries with high malaria burdens (low applicability). In still another example, 394 integrating wild grazing animals on land parcels used for grazing cattle, which are frequently 395 treated for ticks, might increase forage availability for wildlife and reduce tick abundance on land 396 parcels.<sup>95</sup> Tick abundance on land parcels might in turn affect human disease risk, but tick-borne 397 disease incidence in humans has never been measured for this solution (low applicability). As these 398 examples illustrate, many promising solutions had some supporting evidence, but most solutions 399 had notable data gaps (Fig. 5).

400

401 *Optimal solutions achieve goals within resource constraints:* 

In addition to goals (*Impactful* and *Effective*), stakeholders also have resource constraints. Resources determine which solutions can be implemented at the necessary scales and intensities to achieve their desired outcomes (*Affordable, Scalable,* and *Sustainable*) and how big the impact will be for a given resource budget (*Cost-Effective*). Different stakeholders might evaluate these last four criteria differently, because different stakeholders have different goals, priorities, and resources.

408

We note that resource costs and *Affordability* are distinct: cost is the resource price tag, whereas *Affordability* is the ability to pay. For human infectious diseases, public health intervention *Cost-Effectiveness* is usually quantified in disability adjusted life years (DALYs) averted per dollar, but DALYs and cost were rarely reported for the 46 potential solutions reviewed here. This highlights an important area for future research, because for most stakeholders, and perhaps especially those interested in human health outcomes, *Cost-Effectiveness* and *Affordability* will be the most important considerations when choosing a solution.

416

417 For some of the proposed solutions, the potential conservation and health *Impacts* would likely be 418 too small for most stakeholders to justify the cost. For example, invasive python control in the 419 Everglades reduces predation pressures on native vertebrates and might reduce human exposure to the vector-borne Everglades virus.<sup>96</sup> However, python eradication is costly and has not been 420 achieved using existing resources, and maintaining continuous python control efforts at current 421 422 intensities might not be feasible indefinitely (potentially not Sustainable). From a public health 423 perspective, a cheaper and more direct public health or medical intervention might be preferred to 424 python control. However, local stakeholders in the Everglades might value the small human health 425 co-benefits from python control, even if their main goal is a potentially large conservation Impact. 426 As this example illustrates, sometimes a small *Impact* in one sector (health or conservation) can

427 be valued because it accompanies a large *Impact* in another sector, or because it fully addresses a

- 428 small local problem.
- 429

430 It is often difficult to quantify the net value associated with all positive *Impacts* in all sectors for a 431 given intervention. However, identifying these potential *Impacts* explicitly can help stakeholders 432 to compare among multi-sector interventions.<sup>16,21,22</sup> Ultimately, for any given cross-sector 433 problem, collaborations among stakeholders, economists, social scientists, and implementation 434 scientists might be needed to determine which solution is optimal.

- 435
- 435
- 436 *Evidence-based management under uncertainty:*

437 Data-limited solutions that appear safe and feasible could be ideal for immediate research and 438 adaptive implementation. However, strict adaptive implementation requires that multiple 439 interventions are implemented simultaneously and compared, where approaches are subsequently modified according to what works best.<sup>97</sup> This approach is often infeasible in conservation and 440 public health programs,<sup>97</sup> and it might be even more difficult for multi-sector solutions, leaving 441 442 many data gaps unaddressed. When adaptive implementation is not possible, there might be other 443 ways to fill in data gaps via safe implementation, such as by comparing across different programs 444 that all monitor, evaluate, and share their outcomes. For example, multiple programs are improving 445 hygiene or health care for people who work or recreate in great ape conservation areas, which 446 could increase human health and reduce pathogen spillover from humans to apes.<sup>98</sup> Comparing 447 outcomes across these programs might provide new insights for the IUCN's "Best Practice 448 Guidelines for Health Monitoring and Disease Control in Great Ape Populations".<sup>99</sup> As evidence accumulates for this and other solutions, uncertainty will decline. Data gaps will still likely remain 449 450 prominent in the near future, but action despite uncertainty will already be familiar for most public health and conservation practitioners.<sup>9,10</sup> 451

- 452
- 453 *Conclusions*:

454 The growing Planetary Health field emphasizes the links between human well-being and 455 ecosystem integrity, but there has been limited guidance for how to leverage these relationships to 456 implement viable win-win solutions that specifically reduce human infectious disease burdens. Here, we identified 46 such potential win-win solutions. We found that proposed solutions address 457 458 diverse, context-dependent, and dynamic threats with cross-sector interventions that are equally 459 diverse. Many proposed solutions had the potential to be safe and feasible under some predictable 460 contexts, and some were supported by medium- to high-quality evidence of success. Some had the 461 potential for large human health and/or conservation Impacts, such as forest conservation projects 462 and health system strengthening initiatives. Others had relatively minor Impacts but might still be 463 highly valued by local stakeholders. Synergies like these might be pivotal for achieving the soon-

- 464 to-be revised Sustainable Development Goals.<sup>16,21,22,100</sup>
- 465

Though promising, all proposed solutions had some evidence gaps, and collectively, they did not cover all possible health and conservation threats. Evidence regarding conservation and health *Impacts* and intervention *Cost-Effectiveness* and *Affordability* were especially scarce, highlighting priorities for future research. At present, these data gaps within and among solutions complicate decision-making. More evidence will accumulate if stakeholders invest in research, adaptive implementation, and monitoring and evaluation for existing approaches. New solutions will also

472 be imagined, filling in existing gaps among solutions or addressing new problems as they arise.

473 As new solutions and evidence accumulate, the viability criteria described here can be used to 474 compare and update the evidence database for potential solutions, differentiating the solutions that 475 do not work from those that successfully and cost-effectively advance health and conservation.

476

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## Subject-wide Evidence Synthesis

### Phase 1: Systematic Literature Review

Phase 2: 46 Rapid Reviews



## 1.A Search

Searched peer-reviewed literature for studies that hypothesized or measured outcomes for both conservation and human infectious diseases (N=12,270, see PRISMA diagram).

# **1.B Filter**

Filtered literature to 383 studies with at least one proposed win—win solution using human classification and the colandr machine learning algorithm (see PRISMA diagram).

## 1.C Categorize

Combined papers with similar "levers" or examples into collective case studies, creating a list of 46 proposed win–win solutions.



## 2.A Search

Performed individual, targeted searches for peer-reviewed or gray literature relevant to each of the 46 proposed win–win solutions.



# 2.B Summarize

Summarized information relevant to 20 qualitative categories, iteratively returning to the search stage as needed. Summaries were peer reviewed by an expert and revised accordingly.

## 2.C Evaluate

Evaluated available information for each proposed win–win solution, classifying by threat type, evidence for prior success, and viability criteria.

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**Fig. 1.** To find, evaluate, and synthesize evidence from win–win solutions proposed to both reduce human infectious disease burdens and advance conservation goals, we used a subject-wide evidence synthesis. This method first uses a systematic literature review to identify a landscape of possible interventions and then each intervention is explored using an individual, targeted rapid review.<sup>27</sup> Phase 1 involved 10 researchers and Phase 2 involved 25 case study investigators and 44 external expert reviewers.

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Improve health/hygiene to reduce	Conservation 1 hreats & Evidence	Disease Threats & Evidence
anthroponotic spillover to great apes	<b>076 🛣 🚾 🔼</b> 🔤	梁 第 5 📘
deforestation in Borneo	🏍 🋍 <del>*</del>	🕸 💐 🔍 🛛 🕅
Add HIV/AIDS services to conservation	🔉 🔸 👗	۵
Population, Health, Environment programs		* L
for health and conservation	õr 🗰 🗫	梁 <b>9</b>
Improve community reproductive health to improve marine conservation	🛍 🐋 🕅 🕅	<i>₿\$</i>
Control mosquitoes and West Nile virus	<b>*</b>	ä M
transmission to birds and people		
exposure to pathogens		S L
Manage reservoirs for aquatic conservation	<b>8- 1</b> 👌 🛉	
Use poultry biosecurity to reduce influenza		
spillover and wild bird conflicts		<b>黎</b> M
to reduce spillover and conserve bats	🏍 🏥 🛫 🔥 📃	\$
Treat ballast water to control invasive	æ <b>č A</b>	0
species and water-borne pathogens Reduce pesticide use with integrated pest		* L
and vector management	<b>616 A</b>	\$\$ <b>\$</b>
Manage nutrient runoff to reduce eutrophication and water-borne diseases	8% 🛍 🖀 💾	∰∰ \$\$\$ M
Regulate aquaculture antimicrobial use to		
limit pathogen resistance		*
and pathogen invasion	🚾 👃 M	🙊 🂐 M
Ban hunting and consumption of wildlife to	🛧 👗 🗕	<b>Ö</b>
Regulate wildlife trade for pets or		254-264 D
traditional medicine to reduce spillover		楽 後
Use skirts on sap pots to reduce Nipah virus spillover and conserve bats	Sta 🐋	M 🕅
Use clean cookstoves to reduce biomass	<b>.</b>	ă∰ M
burning and respiratory infection		××× ₩
dilute sylvatic disease transmission	🏍 🏥 🐋 🔺 🔺 🔤	¥ L
Forest conservation to reduce malaria risks	26 🖈 👗 🚽	M N
"Re-wild" northeastern North America to		**
control Lyme disease	ot i 🔤 🛣 🔼 🔤 🔤	¥ L
spillover	8% <u>m</u> 🛦 💧 🗖	\$
Conserve or restore forests to prevent		<b>Q</b>
plague outbreaks in Madagascar Introduce mosquito predators to reduce		* L
malaria and pesticide use		梁 <b>《</b>
Conserve or restore wetlands to reduce water-borne pathogen transmission	🛍 🗨 🖀 📕	\$\$ <b>\$</b> \$}
Conserve/restore forests and riparian		10 A A A
buffers to reduce childhood diarrhea Restore native river prawns to control		***
human schistosomiasis	🍋 🖈 📕 📘	k M
Conserve dung beetles to reduce soil- transmitted belminthiases	an 🛍 🖀 🗖	<b>2</b> 2
Control invasive rats to reduce disease and		10 B A A
restore native species		* 35 1
transmission and conflicts with bats	💏 🏙 🐋 📃 📃	M M
Conserve endemic medicinal plants that	₩ the first the first the first tensor tens	Q 2
Conserve leopard to reduce human rabies		<u>米</u>
risks	076 🎹 🔭	<b>梁</b>
Integrate wildlife and cattle grazing and reduce tickborne disease	🂑 🛱 M	\$\$ 🔪 🖿
Restore vulture populations to control		<u>م</u>
human rabies Manage free ranging cats to reduce		*
predation and pathogen transmission	iii 😅 📕	<b>業 9</b> ら
Remove invasive honeysuckle to conserve wildlife and control vectors	â e	🖄 🏶 🚺 🕅
Conserve wild vertebrate communities to		8
dilute Chagas disease risk		
control chronic wasting disease	õ 🔁 📩	۹۸ L
Conserve wetlands or bird biodiversity to	and a market and a market	<b>懲</b>
Use dog tick collars to reduce Rocky		<u>後</u>
Mountain spotted fever transmission	(TR	<b>₩</b>
vaccinate dogs to reduce transmission to wildlife and human rabies	H	🔅 Н
Control pythons to reduce predation and	<b>A</b>	澎
Everglades virus transmission Build bat boxes to conserve bats and reduce		- m
human malaria		۷.
Create nest boxes to conserve raptors and reduce Hanta virus risk	<u></u>	<b>\$</b>
Vaccinate prairie dogs to protect humans	 B¥	Ø
and ferrets from plague	<u>••</u> •	*
<ul> <li>Primary conservation</li> <li>No existing</li> <li>threat</li> <li>evidence</li> </ul>	Low Medium Evidence quality evidence qua	lity High

**Fig 2.** A menu of 46 potential solutions for advancing conservation goals and controlling human infectious diseases. For icon definitions, see Fig. 4.



Fig. 3. Six lose-lose scenarios that could be improved with win-win solutions that reduce 741 742 human infectious disease burdens and advance conservation goals. Health systems that provide 743 affordable healthcare in Indonesian Borneo reduce human disease burdens and illegal logging 744 undertaken to pay for health care. Vector control is a public health intervention that might also 745 benefit biodiversity, as in the case of North American birds susceptible to West Nile virus. Law 746 and policy interventions that ban importation of non-native wildlife reservoirs (e.g., pouch rats) 747 prevent spillover to humans and native wildlife. Education and outreach empower people to live 748 safely with bats, reducing zoonotic spillover risk to people (e.g., Nipah virus) and potentially 749 reducing human-bat conflicts. Species management, like vaccinating or sterilizing free-ranging 750 domestic dog populations, reduces rabies transmission from dogs to humans and African 751 carnivores. Ecosystem management interventions, like restoring wetland vegetation, reduce the 752 survival of human and wildlife pathogens in the environment while restoring wildlife habitat. This 753 figure was commissioned from artist Hiram Henriquez, and all photographs were either used under

- creative commons licenses or purchased with commercial licenses (i.e., iStockphoto). Photographs of bamboo skirts were used with permission from Fernando Garcia and Nazmun Nahar.



757 758 Fig. 4. Solutions are widespread and diverse. (A) The solutions covered all continents (except Antarctica), including countries with both high and low burdens of infectious diseases, as 759 measured by total disability adjusted life years (DALYs) and reported by the World Health 760 Organization for 2016.<sup>101</sup> (B) The solutions also covered most major pathogen taxa (except 761 762 fungi) and transmission modes, where vector-borne transmission was the most common primary 763 transmission mode across solutions. (C) Seven IUCN threat classes were considered the primary 764 conservation threat addressed by at least one solution. In contrast, Transportation, Climate 765 Change, and Energy and Mining were only ever secondary conservation threats addressed by any 766 solution.



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Fig. 5. Viable solutions can be identified and evaluated by 11 criteria. We evaluated five 768 criteria to determine whether solutions were demonstrably Harmless, Contained, Consistent, 769 Feasible, and Acceptable in some predictable contexts given evidence available now, or whether 770 771 they were data deficient for those criteria. Stakeholders can evaluate six other criteria based on 772 their priorities and resource constraints (Impactful, Effective, Affordable, Scalable, Sustainable, 773 and *Cost-Effective*). \*Three potential solutions had evidence for trade-offs that were at present 774 unmitigable or unpredictable and were thus categorized as not Harmless (n=2) and/or not 775 Consistent (n=2). All solutions were categorized as Contained or data deficient for the Contained 776 criterion (not *Contained*: n=0). <sup>†</sup>No potential solutions had evidence for unmediatable barriers to 777 implementation: not *Feasible* (n=0), not *Acceptable* (n=0).

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Fig. 6. Most solutions had evidence gaps. Each cell contains the number of solutions that had a given composite evidence quality score from rubrics based on evidence diversity, consistency, and applicability (None = hypotheses and anecdotes; Low = some supporting studies with moderate to major gaps, inconsistency, or limited applicability; Medium = several lines of evidence that are mostly consistent and applicable; and High = diverse, consistent, and highly applicable evidence leaves little to no uncertainty regarding the outcome). The green color scale is used to emphasize that most solutions were supported by low or medium evidence for conservation and/or health outcomes.