# Aversive medical treatments signal a need for support: a mathematical model Mícheál de Barra<sup>\*†‡1</sup>, Daniel Cownden<sup>\*§2</sup>, and Fredrik Jansson<sup>\*¶3,4</sup> Centre for Culture and Evolution, Brunel University London Centre for Culture and Evolution, Brunel University London Centre for Cultural Evolution, Stockholm University

<sup>4</sup>Division of Applied Mathematics, Mälardalen University

#### Abstract

Ineffective, aversive, and harmful medical treatments are common cross-culturally, his-10 torically and today. Using evolutionary game theory, we develop the following model to 11 explain their persistence. Humans are often incapacitated by illness and injury, and are 12 unusually dependent on care from others during convalescence. However, such caregiv-13 ing is vulnerable to exploitation via illness deception whereby people feign/exaggerate 14 illness in order to gain access to care. Our model demonstrates that aversive treatments 15 can counter-intuitively increase the range of conditions where caregiving is evolution-16 arily viable because only individuals who stand to gain substantially from care will 17 accept the treatment. Thus, contemporary and historical "ineffective" treatments may 18 be solutions to the problem of allocating care to people whose true need is difficult to 19 discern. 20

Keywords: cultural evolution; medical anthropology; sick role; iatrogenic disease;
 evolutionary medicine; cooperation; secondary gain.

Media summary: Model suggests harmful medicine's ability to provide trustable
 evidence of patients' need for care may explain their puzzling persistence.

7

9

<sup>\*</sup>All authors contributed equally.

<sup>&</sup>lt;sup>†</sup>Corresponding author. Centre for Culture and Evolution, Room 110, Marie Jahoda Building, Brunel University London, Uxbridge UB8 3PH, United Kingdom. Email: micheal.debarra@brunel.ac.uk

<sup>&</sup>lt;sup>‡</sup>https://orcid.org/0000-0003-4455-6214

<sup>&</sup>lt;sup>§</sup>https://orcid.org/0000-0002-5348-4841

<sup>¶</sup>https://orcid.org/0000-0001-8357-0276

### 25 1 Introduction

The idea that medicine should be unpleasant and aversive is well rooted in the English 26 language. To take ones' medicine is synonymous with enduring a deserved painful or un-27 pleasant experience. Cheats who are themselves cheated *get a taste of their own medicine*. 28 This reputation is well earned: historical medical treatments were often repugnant, danger-29 ous, taboo breaking or painful. Widespread procedures included ingestion of substances like 30 animal wastes, bird nests, human flesh, as well as poisons, emetics, and diuretics. Surgical 31 procedures like blood-letting, cupping, and the reopening of partially healed wounds were 32 common, as was forced feeding or food and water restrictions (Edgerton, 1992; Miton & 33 Mercier, 2015; Sugg, 2008; Wootton, 2006). Within the history of medicine, the idea that a 34 substantial proportion of pre-20th century western medicines were ineffective or harmful is 35 uncontroversial (Hardy, 2006; Wootton, 2006). 36

The long-term popularity of harmful medicine is surprising given that, all else equal, one might expect individual and social learning processes to be biased against adopting cultural innovations that make life poorer, shorter or more difficult (Boyd, Richerson, & Henrich, 2013). It is also puzzling that these medical treatments should be so *unpleasant/aversive*. Patients who opted for a warm bath over bloodletting sacrificed no therapeutic value. But rather than evolving towards gentle, pleasant, or comforting treatments, the medicine that persisted was invasive, macabre and painful – often theatrically so.

One medical intervention, however, is ancient, common, and unambiguously beneficial: 44 caregiving. Caregiving involves keeping patients comfortable and providing food and other 45 resources and may also entail releasing people from duties and providing for their dependents. 46 Cross-cultural research indicates that this kind of care is both essential and widespread. Some 47 anthropologists have argued that the human life history is premised on access to caregiving 48 (Kaplan, Hill, Lancaster, & Hurtado, 2000; Sugiyama, 2004). In small-scale societies, people 49 are often incapacitated by illness or injury and spend protracted periods of time unable to 50 provide for themselves (Hill & Hurtado, 2009). For example, Sugiyama (2004) reports that 51 90% of Shiwiar – forager-horticulturists in the Ecuadorian Amazon – had spent a fortnight 52 or longer incapacitated. Sixty percent of people fared even worse, spending a month or more 53 unable to forage for themselves or their dependants. Without caregiving, an illness or injury 54 of this duration is fatal. However, when people are disabled by disease, others provide food 55 and other care and take over gardening tasks, sometimes for long periods. About one in 56 seven Americans provide care to a friend or family member who is ill or disabled in a given 57 year (Marks, 1996). A proportion of 2.5% of working hours in the UK are lost to sick leave 58 - institutionalised caregiving – and more than half of these are due to minor illnesses or 59

<sup>60</sup> musculoskeletal illnesses (Comer, 2017).

Caregiving is costly to the carer. Sugiyama (2004) tells how "two informants reported that they jointly maintained [a sick woman's] gardens for three months, but stopped when they could no longer sustain the work". In contemporary Western societies, people involved in long-term caregiving experience poorer health (Vitaliano, Zhang, & Scanlan, 2003) and increased mortality risk (Perkins et al., 2012; Schulz & Beach, 1999), suggesting that caregiving costs remain important even when health insurance and/or public health care provision exist.

From an evolutionary perspective, these costs often constitute a wise investment: helping a sick relative through a period of incapacity can have a substantial effect on their and their offspring's mortality. Hamilton's criterion (Hamilton, 1964) for the evolution of care is frequently met (c < rb, where r is relatedness, b is benefit to the sick, and c is cost to carer). This accords with the cross-cultural and historical research discussed above – caregiving is common and important.

#### 74 1.1 Illness deception

Caregiving, however, is open to exploitation via illness deception. From an evolutionary 75 perspective, the problem is simple: the range of conditions where recipients should request 76 care (rc < b) is much broader than the range of conditions where donors should be willing 77 to grant care (c < rb) (Trivers, 1974, highlights a similar conflict in the context of parental 78 care). If illness were *transparent* – that is, donors could accurately estimate how much the 79 recipient would benefit – then this would be of little consequence. Care could be granted 80 only when it benefited inclusive fitness. However, health status is usually opaque. Many 81 debilitating illnesses leave little visible sign upon the body, for example, back pain, hernia, 82 kidney stones, gallstones, diabetes, Lyme disease, and brucellosis. Conversely, many people 83 with visible aberrations (scarring, rashes, disfigurement) are not in any need of care. Even 84 among people with clear illnesses, it is difficult to estimate how much they will benefit from 85 a given transfer of resources. There is good evidence that people harness this ambiguity in 86 order to access caregiving which the donor would not be willing to offer had they complete 87 information about the recipient's disease state. 88

Hysteria, malingering, factitious disorder, secondary gain, and somatisation disorder are terms used to describe a cluster of related phenomena whereby people assume an ill social state without having a commensurate underlying pathology. They differ in the degree to which they seek release from a specific duty versus the general emotional and practical benefits of caregiving, and in the degree to which the deception is consciously planned and executed versus subconsciously motivated or reinforced. Here we refer to any attempt to feign
or exaggerate illness which may result in access to caregiving as *illness deception*, irrespective
of whether the behaviour is unconsciously or consciously motivated, and irrespective of
whether the scale of the deception is severe or more trivial.

Illness deception is common. In one survey of clinical neuropsychologists, 30% of per-98 sonal injury cases and 33% of disability and worker's compensation cases were judged to 99 "probably" involve malingering or symptom exaggeration (Mittenberg, Patton, Canyock, & 100 Condit, 2002). Several authors have highlighted how fluctuations in illness compensation 101 claims appear unrelated to disease prevalence (Gun, 1990; Nicholson & Martelli, 2007). The 102 introduction of compensation processes is associated with increasing pain reports and reduced 103 treatment effectiveness (Rohling, Binder, & Langhinrichsen-Rohling, 1995) and studies have 104 demonstrated that actors can fool health professionals reasonably easily (Norman, Tugwell, 105 & Feightner, 1982). Illness deception has also been documented in the historical record (see, 106 e.g., Withey, 2013). As several authors have argued (Fabrega, 1997; Finlay & Syal, 2014; 107 Steinkopf, 2015, 2016; Tiokhin, 2016), the fitness benefits associated with care may have 108 acted as a selection pressure on symptom presentation. However, such a selection pressure 109 may not always result in honest displays. 110

#### **111 1.2** Aversive medicine maintains honesty

Caregiving can enhance the inclusive fitness of both donor and recipient, but it is vulnerable 112 to exploitation via illness deception. We propose that decreasing the benefit of caregiving 113 via aversive medical treatments can increase the range of conditions where caregiving is 114 evolutionarily viable. This counter-intuitive proposal can be understood as follows: a fixed 115 reduction in the benefit of care via aversive treatment can shift conditions so that illness 116 deception is no longer viable, allowing caregiving to increase in frequency. These added 117 costs to receiving treatment keep communication honest by allowing caregivers to avoid 118 the problem of distinguishing the ill from the illness deceivers, and by allowing those with 119 hard-to-detect or easily imitated illnesses to credibly request care. 120

This result is possible because truly sick people have much more to gain from caregiving than someone engaged in illness deception. For someone with a significant illness, caregiving can prevent death. For someone with a minor illness or no disease, caregiving provides a lesser benefit, like release from duty or additional food. From an evolutionary perspective, if the aversive treatment (e.g. bloodletting or emetics) is of the appropriate cost, then illness deception will not benefit the individual. Concluding his review of symptoms-assignals, Tiokhin (2016) independently arrives at a similar suggestion noting that if "harsh treatments are painful and timeconsuming, the costs of treatment may not be worth it for those feigning injury."

To better understand the circumstances where aversive treatments can enable caregiving to persist, we develop a mathematical model. Models help to direct our attention to key assumptions, as well as suggest predictions that might be tested in the future.

### 133 **2** Model

We formulate an evolutionary model where individuals reproduce as exually, can be healthy or sick, and where they meet other individuals in random interactions. In these interactions, people have a strategy of whether to ask for help, at a cost to the helper (that causes reduction in fecundity) and a benefit (increasing fecundity) to the recipient, if provided, and whether to provide help when asked. Interactions are assorted, such that relatives meet at a certain frequency.

We use evolutionary game theory with fundamental ideas from invasion analysis (May-140 nard Smith & Price, 1973) to explore the interaction of illness deception, harmful medicine 141 and caregiving. Specifically, we are interested in the conditions under which providing help 142 is a stable strategy and those where it is not. Our main question is whether the range of 143 conditions where helping is evolutionarily viable can be increased through the introduction 144 of aversive medicine, that is, whether there are conditions where the only treatment we can 145 expect is aversive. We will first specify the evolutionary model, then describe the simplifying 146 assumptions, and finally derive conditions for helping and asking strategies to be maintained 147 in the population. For clarity, we keep the model simple, with a minimal set of possible 148 strategies, illustrating the main idea of why aversive treatment can be adaptive, and persist 149 even when only benign caregiving cannot. In the Supplementary material, we expand upon 150 this model with more strategies for what kind of treatment to provide and accept. 151

#### <sup>152</sup> 2.1 Specification and assumptions

An individual encounters the *opportunity* to make fraudulent requests for care (ask for help when healthy) with frequency  $f_{\rm h}$ , and the opportunity to make honest requests for care (ask for help when sick) with frequency  $f_{\rm s}$  (whether an individual will actually make or receive a fraudulent or honest request depends on the strategy of the requester). We assume that these frequencies are set at the population level, that is, they are the same for all individuals. Since every opportunity for an individual to request care when ill is paired with an opportunity for another individual to provide that help (conditional on the request being made), an

Variable	Description
$f_s$	Frequency of opportunity to ask for/give care where recipient is sick
$f_h$	Frequency of opportunity to ask for/give care where recipient is healthy
$b_s$	Benefit of care to sick
$b_h$	Benefit of care to healthy
c	Cost of giving care
r	Degree of relatedness

Table 1: The variables of the model.

individual encounters the potential opportunities to provide help with the same frequencies:  $f_{\rm s}$  to a sick individual and  $f_{\rm h}$  to a healthy individual. When asked for help, an individual does not know whether the requester is sick or healthy.

Providing care entails a cost c. Receiving help gives a benefit  $b_{\rm s}$  if the recipient is sick, and  $b_{\rm h}$  if she is healthy. We assume throughout that the benefit of care when sick is greater than when healthy,  $b_{\rm s} > b_{\rm h}$ .

Finally, we assume that there is an assortative mechanism that produces a degree of relatedness r between interacting individuals. Relatedness is here defined as the probability that an allele sampled from the actor will be identical by descent to an allele sampled from the recipient, and hence they will employ the same strategy. We return to this assumption below. The variables of the model are summarised in Table 1.

To derive conditions for *helping* to be maintained in this model, the general idea is 171 to consider the situations where there is a resident strategy at dynamical equilibrium and 172 evaluate the initial growth rate of a mutant strategy in such an environment, the invasion 173 fitness (see e.g. Brännström, Johansson, & Festenberg, 2013). The success of the mutant 174 strategy is then inferred by the growth rate when rare. As is common in invasion analysis, 175 we incorporate the simplifying assumptions that the strategies interact within an infinite 176 monomorphic population, that reproduction is asexual, and that interaction occurs between 177 pairs of strategies. Although these behaviours are cultural traits, our model focuses on the 178 genetic fitness of people who engage in these behaviours. Later, we discuss how this genetic 179 fitness might translate into cultural success. 180

Returning to the degree of relatedness r, suppose that there is a behaviourally relevant allele that causes reduction in personal fecundity c (for cost) while at the same time causing the fecundity of some other individuals to be increased by b (for benefit). Hamilton (1963) showed that in the case of discrete, non-overlapping generations, this allele for a helping behaviour can spread provided that there is some assortative mechanism whereby individuals are more likely to interact with relatives. Specifically, helping behaviour will be favoured by natural selection precisely when rb > c. While this seems relatively straightforward, it should

be noted that many of the plausible assortative mechanisms that might cause interactants 188 to be related, for example spatial structure coupled with limited offspring dispersal, can 189 also serve to localise competition so that the benefits of cooperation are squandered in 190 subsequent competition between relatives (West, Pen, & Griffin, 2002). Here we assume 191 that competition is homogeneous throughout the population, such that competition is not 192 stronger among relatives than among non-kin. This is a simplification, but since our aim 193 is to illustrate the evolutionary potential of aversive medicine rather than to derive exact 194 conditions for values of parameters (that lack empirical data), simplicity and clarity are more 195 important. 196

Relatedness r is thus an input parameter to the model, and is the same throughout 197 the population (as in the signalling model by Maynard Smith, 1991; this is a first-order 198 approximation for frequency change, or a "weak selection" assumption, as described by 199 Rousset & Billiard, 2000). In a scenario with several strategies in the population, this could 200 potentially have a large impact on the dynamics, if we can expect that the success of different 201 strategies will influence r. In our analysis, however, we compare the fitness of residents with 202 the same strategy only to mutants with another strategy, similar to the approach taken by 203 Taylor and Frank (1996), where r remains the same for a rare mutant (see also Gardner & 204 West, 2006, on the relative merits of approaches with closed models where r is determined 205 by demographic assumptions versus open models where it is allowed to vary independently). 206 In fact, as will be obvious in the invasion analysis, r is only relevant in the fitness equation 207 for the mutant, so in our analysis, r can be interpreted as the frequency with which mutants 208 interact with individuals identical by descent, and 1 - r as the frequency with which they 209 interact with the rest of the population. 210

As mentioned earlier, while the person requesting care knows whether they are healthy 211 or sick, the person receiving the request does not (health status is opaque). The possible 212 strategies in this game are thus composed of three components: 1) whether or not to request 213 care when ill, 2) whether or not to request care when healthy and 3) whether or not to 214 provide care when it is requested. This means that there are eight  $(2^3)$  possible strategies, 215 allowing for all possible combinations of the component parts of the strategies. However, 216 three of these strategies weakly dominate the rest and so we limit our analysis to these three. 217 The three dominant strategies are *Deceptive Nonhelper*, which will request care both when 218 ill and healthy and does not provide care when asked, *Honest Helper*, which requests care 219 only when truly ill and provides care when asked, and *Deceptive Helper*, which requests care 220 both when ill and healthy and provides care when asked. Since there is only one non-helping 221 strategy, and no honest non-helpers, we will henceforth refer to *Deceptive Nonhelper* simply 222 as Nonhelper. (Weak domination means that any strategy outside of the set of Nonhelper, 223

Honest Helper and Deceptive Helper can only ever do as well as, but never better, than one of these dominating strategies, regardless of the population profile. All three dominant strategies ask for help when sick. The strategies that are dominated are Honest Nonhelper and the corresponding strategies to the Honest Nonhelper, and the three dominant ones that do not ask for help when sick.)

Let  $\delta$  be the indicator function, that is,  $\delta(x) = 1$  if x is true and  $\delta(x) = 0$  if x is false. The general equation for any of these strategies is

fitness increment = expected benefit of care when sick

– expected cost of caring for sick

+ expected benefit of care when healthy

- expected cost of caring for healthy

- $= f_{\rm s} b_{\rm s} \cdot \delta(\text{ask for care when sick}) P(\text{receive care})$ 
  - $-f_{\rm s}c \cdot \delta(\text{provide care})P(\text{be asked for care by sick})$

 $+ f_{\rm h}b_{\rm h} \cdot \delta(\text{ask for care when healthy})P(\text{receive care})$ 

 $-f_{\rm h}c \cdot \delta(\text{provide care})P(\text{be asked for care by healthy})$ 

- $= f_{\rm s} b_{\rm s} \cdot P(\text{receive care})$ 
  - $-f_{\rm s}c \cdot \delta(\text{provide care})$
  - $+ f_{\rm h}b_{\rm h} \cdot \delta(\text{ask for care when healthy})P(\text{receive care})$
  - $-f_{\rm h}c \cdot \delta(\text{provide care})P(\text{be asked for care by healthy})$

In each of these expressions, fitness is computed as the expected benefit or cost in the 229 following four situations: asking for help as a sick person (the opportunity of which occurs 230 with frequency  $f_s$  and provides a benefit  $b_s$  with probability P(receive care) since all strategies 231 ask when sick); being asked for help by a sick person (with frequency  $f_s$ , providing a cost 232 c if the strategy provides care); having the opportunity to ask for help as a healthy person 233 (with frequency  $f_{\rm h}$ , providing a benefit  $b_{\rm h}$  with probability P(receive care) if the strategy 234 asks for help when healthy); and potentially being asked for help by a healthy person (with 235 frequency  $f_{\rm h}$ , providing a cost c if the strategy provides care and the recipient asks for it 236 when healthy). We examine the fitness expressions for each strategy in turn. 237

We let  $P_{\rm N}$ ,  $P_{\rm H}$  and  $P_{\rm D}$  denote the proportions in the population of *(Deceptive) Nonhelper*, Honest Helper and Deceptive Helper strategies, respectively, and let  $W_{\rm N}$ ,  $W_{\rm H}$  and  $W_{\rm D}$  denote their respective fitness. Then we can compute the fitness benefit for *Nonhelpers* as

$$\Delta W_{\rm N} = f_{\rm s} b_{\rm s} (1-r) (P_{\rm H} + P_{\rm D})$$
$$- f_{\rm s} 0$$
$$+ f_{\rm h} b_{\rm h} (1-r) (P_{\rm H} + P_{\rm D})$$
$$- f_{\rm h} 0$$

A Nonhelper always asks for help, but will receive it only when asking a non-relative (since relatives employ the same strategy, and thus never help), which occurs with probability 1 - r, who employs one of the helping strategies, which occurs with probability ( $P_{\rm H} + P_{\rm D}$ ). A Nonhelper never provides care.

The fitness benefit for *Honest Helpers* is

$$\Delta W_{\rm H} = f_{\rm s} b_{\rm s} (r + (1 - r)(P_{\rm H} + P_{\rm D}))$$
  
-  $f_{\rm s} c$   
+  $f_{\rm h} 0$   
-  $f_{\rm h} c (1 - r)(P_{\rm N} + P_{\rm D})$ 

An Honest Helper will be helped when sick also by a relative, increasing the probability of receiving care when sick to  $r + (1 - r)(P_{\rm H} + P_{\rm D})$ , while she never asks for help when healthy. An Honest Helper provides care when asked (and will always be asked if the recipient is sick). If the recipient is healthy, only a non-relative who employs one of the always asking strategies will use the opportunity to ask for health, which occurs with probability  $(1 - r)(P_{\rm N} + P_{\rm D})$ .

Finally, for *Deceptive Helpers*, we have

$$\Delta W_{\rm D} = f_{\rm s} b_{\rm s} (r + (1 - r)(P_{\rm H} + P_{\rm D}))$$
  
-  $f_{\rm s} c$   
+  $f_{\rm h} b_{\rm h} (r + (1 - r)(P_{\rm H} + P_{\rm D}))$   
-  $f_{\rm h} c (r + (1 - r)(P_{\rm N} + P_{\rm D}))$ 

A Deceptive Helper is not different to an Honest Helper when sick. Given the opportunity, a Deceptive Helper will ask for help also when healthy, and has the same probability to receive it as when sick. Contrasting to Honest Helper, a Deceptive Helper will be asked for help by a healthy relative, increasing the probability of providing care for healthy to  $r + (1 - r)(P_{\rm N} + P_{\rm D})$ .

$Mutant \ \backslash \ Resident$	Nonhelper	Honest Helper	Deceptive Helper
Nonhelper		$\frac{b_{\rm h}}{c} > \frac{\frac{f_{\rm s}}{f_{\rm h}}}{1-r} \left( r \frac{b_{\rm s}}{c} - 1 \right)$	$\frac{b_{\rm h}}{c} < \frac{1}{r} \left( 1 + \frac{f_{\rm s}}{f_{\rm h}} - \frac{f_{\rm s}}{f_{\rm h}} \frac{b_{\rm s}}{c} \right)$
Honest Helper	$\frac{b_{\rm s}}{c} > \frac{f_{\rm h}}{f_{\rm s}} \left(\frac{1}{r} - 1\right) + \frac{1}{r}$		$\frac{b_{\mathrm{h}}}{c} < r$
Deceptive Helper	$\left  \frac{b_{\rm h}}{c} > \frac{1}{r} \left( 1 + \frac{f_{\rm s}}{f_{\rm h}} - \frac{f_{\rm s}}{f_{\rm h}} \frac{b_{\rm s}}{c} \right) \right $	$\frac{b_{\rm h}}{c} > r$	

Table 2: Conditions for when mutant can invade single resident strategy.

### <sup>252</sup> 2.2 Evolutionarily stable strategies

Using these expressions for fitness we can investigate the conditions under which each pure strategy is resistant to invasion from rare mutants of the other strategies, that is, the *evolutionarily stable strategies* (ESS) (Maynard Smith & Price, 1973).

The invasion conditions are derived in the supplementary materials, and are summarized in Table 2.

The outcomes of these conditions can be visualized as a map in a parameter space. In the plots that follow, relatedness has been fixed at r = 0.25 and the frequencies have been fixed at  $f_s = f_h = 0.25$ , and we plot using the normalised parameters  $\frac{b_h}{c}$  and  $\frac{b_s}{c}$ , since it is the ratio of cost-to-benefit that determines the evolutionary outcomes, not the absolute values (see also Table 2). In the supplementary material, we show that qualitatively similar results hold for a range of r,  $f_s$  and  $f_h$  values.

In one of the limiting cases, when  $c < rb_{\rm h}$  (which implies that  $c < rb_{\rm s}$  since we assume  $b_{\rm h} < b_{\rm s}$ ), we have from Hamilton's rule that *Deceptive Helper* is the only ESS, in the other extreme case where  $c > rb_{\rm s}$  (which again implies that  $c > rb_{\rm h}$ ) Hamilton's rule gives (*Deceptive*) *Nonhelper* as the only ESS. The interesting cases are thus in the parameter range where  $rb_{\rm h} < c < rb_{\rm s}$ , that is, where helping the ill is evolutionarily viable, but helping the healthy is not.

As can be seen in Figure 1, there is a broad range of conditions where the possibility of 270 deception undermines caregiving (i.e., where *Nonhelpers* can establish – the light regions, 271 mainly the orange and yellow regions, where they cannot be invaded, and to a lesser extent 272 the dark orange region, where all strategies can be invaded). Now we consider the potential 273 impact of aversive treatments on these evolutionary outcomes. For simplicity we assume 274 that an aversive treatment reduces the benefit of receiving care when healthy and when 275 sick in equal measure. Under this assumption, and in the context of Figures 1 and 2, the 276 introduction of an aversive treatment can be thought of as shifting a model's point in the 277 parameter space downward and to the left at a 45 degree angle (i.e., to follow a straight 278 line with slope 1, in the left direction, where the length of the shift is determined by the 279 aversiveness of the treatment). Figure 2 highlights those regions in the original parameter 280

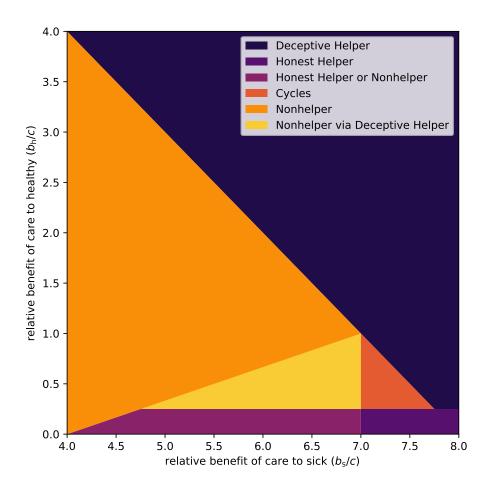


Figure 1: Evolutionarily stable strategies when relative benefit to sick  $(b_s/c)$  and relative benefit to healthy  $(b_h/c)$  vary. Relatedness is set to r = 0.25, and opportunities for illness deception and legitimate care request occur with equal probability,  $f_s = f_h = 0.25$ . Colours depict which pure strategies are stable for a given pair of benefits, for which they can resist invasion. The dark, blue/purple regions, are where helping strategies can be maintained: in the blue top right region *Deceptive Helper* dominates; in the purple bottom right region *Honest Helper* dominates; and in the violet bottom left region *Honest Helper* and *Nonhelper* are in a stalemate situation where both are evolutionarily stable, with neither being able to invade the other. Helping is not maintained in the bright, yellow/orange regions: in the orange leftmost region *Nonhelper* dominates; in the yellow central left triangular region, the dominance of *Nonhelper* is a direct result of the *Deceptive Helper* strategy being able to invade the *Honest Helper* strategy, paving the way for an invasion by Nonhelpers; and in the red central right triangular region no strategy dominates, with *Honest Helper* being able to invade *Nonhelper*, which in turn is able to invade *Deceptive Helper*, which is in turn able to invade *Honest Helper*, and so on in a cycle.

space where caregiving is undermined by illness deception (i.e., the light yellow/orange/red regions, where *Nonhelpers* can establish), but where it is possible for caregiving to become an ESS, via a judicious choice of the degree of aversiveness of the treatment. Figure A.1 in the supplementary material shows that to the extent that illness deception undermines caregiving, aversive medicine can help prevent this erosion. That is to say, aversive medicine plays a more important role when illness deception is common.

In this model, we compare a universe where caregiving is benign and has no side effects 287 to one where it is aversive, showing that treatment can become common where in the former 288 universe it would not. However, the model does not allow for alternative practices to compete 289 directly, and for caregiving and accepting treatment to be contingent on accepting aversive 290 treatment when treatment without side effects may be a viable option. In the Supplementary 291 material, we extend the model to see whether aversive treatment can be sustained also in 292 direct competition from beingn treatment without side effects. Such a model significantly 293 expands the number of possible strategies and makes the model less perspicuous, but, con-294 sidering the same parameter space as in Figures 1 and 2, the results can be summarised as: 295 (1) there will only be beingn treatment where Deceptive Helpers constituted an ESS, but 296 (2) the parameter space in which caregiving becomes possible due to aversive treatment (the 297 dotted and dashed regions) expands. 298

The following explicit empirical predictions are based on the original model, but the general qualitative predictions are consistent also with the extended model.

### **3 Empirical predictions**

Here we outline how the theory and model generate predictions both about *when* we would expect to see harmful medicines and *how harmful* we would expect them to be. We also discuss how existing findings relate to these predictions and speculate on how they might be tested in the future.

#### <sup>306</sup> 3.1 No care without aversive treatment

If a function of medical treatment is to legitimise one's request for care, then people should be less inclined to provide care to those who do not undergo treatment, since if potential illness deceivers could access care without aversive treatment, then treatment's capacity to stabilise caregiving would evaporate. Thus, one prediction from our theory is that care will often be conditional on the acceptance of a treatment.

This prediction is consistent with Parsons' sociological analysis of the *sick role* (1951).

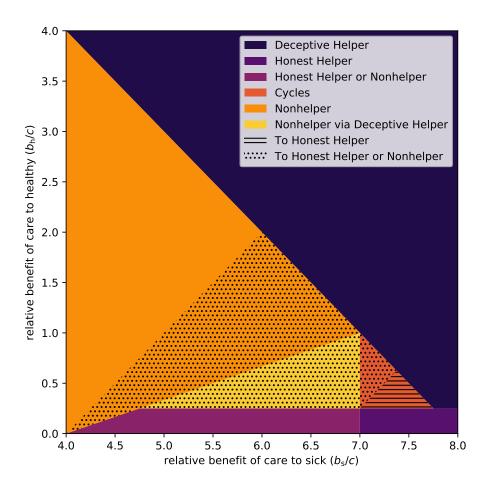


Figure 2: Evolutionarily stable strategies and potential helper ESSs when a harmful medical treatment is introduced. Relatedness is set to r = 0.25, and opportunities for illness deception and legitimate care request occur with equal probability,  $f_s = f_h = 0.25$ . The dotted region is where the dominance of *Nonhelper* (orange and yellow areas) can be eroded or cycling between *Nonhelper* and the other strategies (red area) can be stopped by aversive medicine, creating a stalemate situation where both *Nonhelper* and *Honest Helper* are evolutionarily stable. The lined area shows where aversive treatments can stop the cycling of strategies and make *Honest Helper* the sole ESS.

When someone occupies a sick role, they are released from their social obligations and not 313 held morally responsible for the additional burden that this places on others, but, crucially, 314 they must do everything possible to exit the sick role, including taking any medications 315 or treatments recommended by medical professionals. The theory outlined here suggests 316 that this obligation to undergo the trials of treatment helps to maintain the stability of 317 the institution in the face of would-be deceivers. Indeed, Parsons and Fox (1952) note how 318 negative aspects of interaction with the health care system "are the penalties which give 319 impetus to the patient's desire to re-achieve wellness" 320

Qualitative research suggests that some patients have noted that access to care seems to be a function of treatment acceptance. A study of chronic pain sufferering (Kleinman, 1988, quoted by Glenton, 2003) reports that:

The surgeries have had one clearly positive effect, in Howie's view. They have 324 created icons of his travail, scars that he can show people, that he can touch 325 himself to assure himself that there is something 'physically wrong' with his 326 back. After each of his surgeries, he felt that his family, fellow police officers, 327 and doctors became more sympathetic. As he contemplates yet another major 328 surgical procedure, this latent social function of surgery is a large part of the 329 decision making, since his overall judgement about the surgeries is that they 330 have made things worse. 331

Data on social support provided to patients who are randomised to undergo invasive or non-invasive treatments in clinical trials would provide an interesting test: if patient's need for care is opaque, we would predict greater social support for patients who undergo the invasive procedure.

### 336 3.2 How harmful should treatments be?

In order to prevent illness deception from undermining caregiving, conditions must be set such that *Nonhelper* cannot invade. Table 2 shows that aversive medicine could potentially prevent *Nonhelper* from invading *Honest Helper*, but not from invading *Deceptive Helper* (since reducing  $\frac{b_h}{c}$  and  $\frac{b_s}{c}$  would decrease the left-hand side and increase the right-hand side of the inequality). However, some amount of harm could prevent *Deceptive Helper* from invading *Honest Helper*. Thus, there are two regions where aversive medicine could maintain caregiving: it can stop *Nonhelpers* from invading directly if

$$\frac{b_{\rm h}}{c} - a \le \frac{f_{\rm s}/f_{\rm h}}{1 - r} \left( r \left( \frac{b_{\rm s}}{c} - a \right) - 1 \right)$$

<sup>344</sup> and from invading by way of *Deceptive Helper* if

$$\frac{b_{\rm h}}{c} - a \leq r$$

where  $a \ge 0$  is the amount of harm of an aversive treatment. The smallest amount of a that will meet the inequalities is thus

$$a = \begin{cases} \frac{b_{\rm h}}{c} - r & \text{if Nonhelpers could invade only through Deceptive Helper} \\ \frac{\frac{b_{\rm h}}{c} - \frac{f_{\rm s}/f_{\rm h}}{1 - r} \left(r \frac{b_{\rm s}}{c} - 1\right)}{1 - \frac{f_{\rm s}/f_{\rm h}}{1 - r} r} & \text{if Nonhelpers could invade directly} \end{cases}$$

Note that the denominator in the second expression is negative if (and only if)  $\frac{r}{1-r} > \frac{f_{\rm h}}{f_{\rm s}}$ , that is, when there are few opportunities to ask for help when healthy as compared to when sick, and/or most of these requests will be to relatives. For example, if  $f_{\rm s} \leq f_{\rm h}$  and r < 0.5, then the denominator is positive.

The first expression increases with  $\frac{b_h}{c}$ , and so does the second when the denominator 351 is positive, so we would expect to see more severe treatments when the benefits of illness 352 deception are great. Therefore we predict that treatments will be more harmful, for example, 353 in times of intergroup conflict among potential combatants relative to times of peace or 354 among non-potential combatants. Illness deception has long been a problem for armed 355 forces. The problem became acute in the first world war due to uncertainty over whether 356 neuropsychological problems like "shell shock" were instances of illness deception. Wessely 357 (2003) notes that the growing suspicion among military and medical elites, coupled with a 358 shortage of men, meant that "German (and of course British) treatments for the war neuroses 359 became increasingly punitive". 360

A related prediction is that treatments should be more harmful in societies where people engage in dangerous foraging activities (e.g., hunting large mammals) than in societies where resource acquisition is safer: this hypothesis may be testable with cross-cultural ethnographic datasets. The idea that care is contingent on harsher treatment when the benefits of access to the sick role is higher might also be tested in vignette experiments.

On the other hand, to prevent *Deceptive Helper* from invading, we expect medicines to be less harmful when the denominator is large, that is, when the costs to the caregiver are substantial. This is somewhat counter-intuitive – would not a stronger deterrent be preferable when the costs of caregiving are large? – but it can be understood as a consequence of the fact that the cost of caregiving is disproportionately borne by relatives. Hence, from an inclusive fitness perspective, the costs to relatives of a request for care will not outweigh the benefits to self from that care. Finally, the closer the relatives, the more benign the treatment <sup>373</sup> to stop *Deceptive Helpers*. We know of no data that test these predictions directly.

In the region where *Nonhelpers* can invade *Honest Helpers* directly, the aversiveness of medicine can either increase or decrease with costs and high relatedness, depending on the other variables. For example, if

$$\frac{f_{\rm s}}{f_{\rm h}} \frac{r}{1-r} b_{\rm s} > b_{\rm h}$$

then medicines should be more harmful when the costs to the caregiver are large, while in the opposite case, they are expected to be less harmful. We refrain from going into further detail, the main point being that predictions are more complex when *Nonhelpers* can invade directly.

#### <sup>381</sup> 3.3 When should harmful medicine be more common?

Aversive medical treatments are expected to be more common when illness deception is possible. In situations where need is largely transparent – for example, when diagnostics are reliable, where the disease has obvious, familiar causes, or illness that is difficult to fake – then costly treatments are not needed. Epidemic infectious diseases that infect large numbers of people and that have consistent symptomatology and consequences will negate the need for harmful treatments. So will the reliable diagnostic and prognostic methods that become common over recent decades.

The quotation above from the backpain sufferer (Kleinman, 1988) illustrates the partic-389 ular importance of visible and significant treatment when the visible symptoms are absent. 390 Similarly, in her study of illness behaviour in Fiji, Trnka (2007) finds that women whose need 391 for care is opaque seek costly legitimation of their problem via written doctors' prescriptions, 392 even though the medication they desire is widely available. We might predict that individ-393 uals with, for example, an obvious cut rather than non-obvious muscle injury would be less 394 concerned about this prescription. A related prediction amenable to laboratory testing is 395 that acceptance of costly treatment should be less relevant to would be caregivers when the 396 need is transparent. 397

Generally speaking, cultures which deploy aversive treatments for ailments where the 398 costs of providing and the benefits of receiving care, corresponding to those in the hatched 399 parameter space of Figure 2, are less likely to have their caregiving practices undermined 400 by illness deception. This has a number of implications. When c is very low, caregivers 401 have little to lose and much to gain by offering care freely. As c increases, we would expect 402 costly treatments to become more common (until c becomes so substantial that the benefit to 403 healthy scaled by costs to provider is close to zero, in which case costly treatments are again 404 not needed); in Figure 2, this is equivalent to a move from the top right diagonally down 405

and left into the hatched central area. In the real world a range of factors will influence  $c_{i}$ 406 including the caregiver's time or energy, food availability, or the scale of the care requested. 407 It follows that childhood illnesses are less likely to be treated with harmful medicines. 408 Even when healthy, children's economic contribution is limited and hence the loss of their 409 labour is a less significant problem. Moreover, children require substantial care independent 410 of illness. Thus c will generally be relatively low. For similar reasons, the elderly and infirm 411 are also less likely to be treated with harmful medicines. Although the perceived absence 412 of side-effects is an important reason that children are given complementary and alternative 413 medicines (Cuzzolin et al., 2003), other datasets are needed to test this prediction more 414 directly. 415

### 416 4 Discussion

Our model suggests that the judicious introduction of harmful treatments, in conjunction 417 with effective caregiving, broadens the range of conditions where caregiving is evolutionarily 418 viable. There is a broad range of conditions, that is, relative cost-benefit ratios of receiving 419 and providing care, where the possibility of illness deception renders caregiving evolutionar-420 ily inviable. We show that the introduction of aversive medicine that reduces the benefit of 421 receiving care can in some cases transform the underlying strategic situation so that caregiv-422 ing becomes evolutionarily viable where previously it was not. This is possible because the 423 benefit of care for the truly sick is greater than the benefit of care for the illness deceiver. 424 The model shows that there is scope for the benefit of care to be reduced for both the ill 425 and the illness deceivers in such a way that illness deception is no longer the evolutionarily 426 dominant strategy, allowing caregiving the chance to increase in frequency. 427

Note that the current model has no bearing on the spread of *beneficial* or *effective* treat-428 ments. Treatment benefit and harm are orthogonal – a single treatment can be both very 429 aversive and very helpful (e.g., surgery). We suggest that selection pressures sometimes 430 favour treatments higher in the harm dimension, but it is plausible that other selection 431 pressures may favour treatments higher in the benefit dimension, particularly because the 432 benefit of effective treatments is often only realised if the recipient is truly ill. Some specific 433 theoretical work as well as general cultural evolution models suggest that treatments may 434 also evolve towards helpfulness (Henrich & Henrich, 2010; Tanaka, Kendal, & Laland, 2009). 435 Although the model above analyses treatments as if they were genetic traits, medicine is 436

<sup>430</sup> largely a cultural phenomenon. However, there are several processes by which genetic fitness
<sup>437</sup> could translate into cultural fitness. Once a harmful medical practice emerges in a commu<sup>439</sup> nity, people who accept or demand the use of this signal will, on average, have better health

than those who reject it. Better health translates into more, healthier, children, and thus if 440 medical beliefs are passed from parent to child, its frequency will increase within the group. 441 Moreover, people are probably more inclined to learn from healthy peers and parent/elders 442 than from the ill. Thus oblique and horizontal transmission may also facilitate trait spread. 443 Alternatively, if healthy individuals are better transmitters of cultural practices of medicine 444 and helping behaviour, and transmission takes place in the same assorted interactions as 445 the opportunities for help, then our model translates into a cultural evolution model, with 446 fitness being a measure of cultural transmission from an individual. Another possibility is 447 that as a result of individual learning, or cultural or genetic evolution, human cognition is 448 generally sensitive to the risk of deception (including illness deception) as well as to cues 449 (such as treatment acceptance) that indicate such deception is unlikely. Such a psychology 450 would provide fertile ground for the cultural evolution of harmful therapies. 451

The value of harmful medicine is not dependent on people understanding its functional 452 role. We suggest that over many generations, harmful medicine spreads within a community 453 because people who use it end up healthier (and having healthier kin) than people who do 454 not. "Deterrent" medicine may work better when its true function is hidden, since if the 455 message component were obvious, then skilled illness deceivers might circumvent the treat-456 ment through persuasion or appeals to other kinds of evidence that purport to demonstrate 457 their illness. Those who suspect illness deception would need to make an explicit accusation. 458 an act likely to damage relationships, whether or not illness deception is taking place. 459

There are parallels between the processes described here and costly signalling theory 460 (Grafen, 1990; Zahavi, 1975). However, in many costly signalling contexts, what varies is 461 the costs that it takes to produce a given signal. In the present case, the cost of producing 462 the signal is similar across all individuals. What varies is instead the benefit that results from 463 the production of this signal such that people who are sicker stand to gain much more from 464 a unit of care than people who are less sick. Hence the fixed cost of aversive treatments will 465 deter all but people who stand to gain substantially. The chick begging model developed by 466 Godfray (1991) has a similar dynamic. Chicks pay a cost to request food through begging, 467 and the benefit of a unit of that food is lower if the chick has been recently fed. Like in the 468 medical case outlined here, the fixed costs of requesting enable donors to efficiently identify 469 situations where that transfer of resources is most useful. Unlike in the medical case, the 470 transfer of resources is unidirectional, from parent to offspring. 471

While we have built this model within a kin selection framework, other processes may enable the evolution of caregiving as well as illness deception and harmful medical treatments. According to direct reciprocity theory (Trivers, 1971), individuals will provide each other with care in times of need with the expectation that this care is reciprocated in the future.

Care based on such reciprocity is less subject to erosion via illness deception, since the carer's 476 fitness is enhanced by the return of care when they fall ill in the future. Thus, whether the 477 care benefits the requester a lot (if they are truly sick) or a little (if they are engaging in 478 deception) is of little consequence to the carer; they should only be concerned about the 479 availability of care to themselves in the future. However, direct reciprocity depends on a 480 predictability and symmetry of illness or injury that may be rare in nature, since people have 481 cannot predict if, when, and how much care they will need in the future (see also Clutton-482 Brock, 2009; Raihani & Bshary, 2011). Also, if they predict that the requester may never 483 recover to a degree that would enable them to return the care, direct reciprocity alone will 484 not sustain caregiving. 485

Indirect reciprocity (Nowak & Sigmund, 1998), in which people with a reputation as 486 caring are then cared for if they request it, may be more likely to sustain caregiving. Such 487 a reputation-based system depends less on a symmetry of need between partners and the 488 predictability of illness or injury. However, in a society where people are inclined to provide 489 care so as to maintain a caring reputation, an incentive to engage in illness deception will 490 exist. Since the amount of care available within this society is finite, frequent illness deception 491 will diminish the care available to people with true illnesses. Thus, a sort of tragedy of the 492 commons may result, whereby illness deception reduces the care available to the truly ill, 493 who benefit much more from each unit of care. However, like in the kin selection model we 494 developed here, harmful treatments that impose a fixed cost on every requester will diminish 495 this problem, since only people who stand to benefit substantially from care will request it in 496 the face of these costs. Moreover, aversive treatments may enable ill actors who require care 497 to maintain an "honest" reputation; this may be important for maintaining or developing 498 new cooperative relationships in contexts where partner selection occurs (Barclay, 2013; 499 Baumard, André, & Sperber, 2013). 500

In conclusion, the theory presented here suggests an explanation for several puzzling 501 questions of medical cultural evolution and contributes to a growing literature on the evolu-502 tion of medical practice (De Barra, 2017; De Barra, Eriksson, & Strimling, 2014; Jiménez, 503 Stubbersfield, & Tehrani, 2018; Miton & Mercier, 2015; Miton, Claidière, & Mercier, 2015; 504 Steinkopf, 2017; Tanaka et al., 2009). Medicines may serve not just to cure disease but also 505 to deter illness deception. Many treatments that are directly harmful may be indirectly ben-506 eficial in that they help to expand the range of circumstances in which valuable caregiving 507 can persist. 508

### 509 5 Required statements

Acknowledgements We thank Magnus Enquist for valuable discussions and Marijn de Bruin for his comments on a draft of this manuscript.

<sup>512</sup> Author Contributions The authors made an equal contribution. MdB developed the <sup>513</sup> idea. DC and FJ developed the model. MdB, DC, and FJ wrote the manuscript.

Financial Support MdB was supported by the Roy Weir Career Development Fellowship.
FJ was supported by the Knut and Alice Wallenberg Foundation (grant number 2015.0005).

<sup>516</sup> **Publishing Ethics** This study involved no participants.

517 Conflict of Interest There are no relevant conflicts of interest.

### **List of Figures**

1 Evolutionarily stable strategies when relative benefit to sick  $(b_s/c)$  and relative 519 benefit to healthy  $(b_{\rm h}/c)$  vary. Relatedness is set to r = 0.25, and opportunities 520 for illness deception and legitimate care request occur with equal probability, 521  $f_{\rm s} = f_{\rm h} = 0.25$ . Colours depict which pure strategies are stable for a given pair 522 of benefits, for which they can resist invasion. The dark, blue/purple regions, 523 are where helping strategies can be maintained: in the blue top right region 524 Deceptive Helper dominates; in the purple bottom right region Honest Helper 525 dominates; and in the violet bottom left region Honest Helper and Nonhelper 526 are in a stalemate situation where both are evolutionarily stable, with neither 527 being able to invade the other. Helping is not maintained in the bright, 528 vellow/orange regions: in the orange leftmost region *Nonhelper* dominates; in 529 the yellow central left triangular region, the dominance of Nonhelper is a direct 530 result of the *Deceptive Helper* strategy being able to invade the *Honest Helper* 531 strategy, paving the way for an invasion by Nonhelpers; and in the red central 532 right triangular region no strategy dominates, with *Honest Helper* being able 533 to invade *Nonhelper*, which in turn is able to invade *Deceptive Helper*, which 534 is in turn able to invade *Honest Helper*, and so on in a cycle. . . . . . . . . . 535

10

2Evolutionarily stable strategies and potential helper ESSs when a harmful 536 medical treatment is introduced. Relatedness is set to r = 0.25, and op-537 portunities for illness deception and legitimate care request occur with equal 538 probability,  $f_{\rm s} = f_{\rm h} = 0.25$ . The dotted region is where the dominance of 539 Nonhelper (orange and yellow areas) can be eroded or cycling between Non-540 *helper* and the other strategies (red area) can be stopped by aversive medicine, 541 creating a stalemate situation where both Nonhelper and Honest Helper are 542 evolutionarily stable. The lined area shows where aversive treatments can 543 stop the cycling of strategies and make *Honest Helper* the sole ESS. . . . . . 544

## 545 6 References

Barclay, P. (2013). Strategies for cooperation in biological markets, especially for humans.
 *Evolution and Human Behavior*, 34 (3), 164–175.

12

- Baumard, N., André, J.-B., & Sperber, D. (2013). A mutualistic approach to morality: The
  evolution of fairness by partner choice. *Behavioral and Brain Sciences*, 36(1), 59–78.
- <sup>550</sup> Boyd, R., Richerson, P. J., & Henrich, J. (2013). The cultural evolution of technology: Facts
  <sup>551</sup> and theories. In P. J. Richerson & M. H. Christiansen (Eds.), *Cultural evolution: Society,*<sup>552</sup> language, and religion (pp. 119–142).
- Brännström, Å., Johansson, J., & Festenberg, N. von. (2013). The Hitchhiker's Guide to
  Adaptive Dynamics. *Games*, 4(3), 304–328.
- <sup>555</sup> Clutton-Brock, T. (2009). Cooperation between non-kin in animal societies. *Nature*, 462(7269),
   <sup>556</sup> 51.
- <sup>557</sup> Comer, M. (2017). Sickness absence in the labour market: 2016. Office of National Statistics.
- <sup>558</sup> Cuzzolin, L., Zaffani, S., Murgia, V., Gangemi, M., Meneghelli, G., Chiamenti, G., & Benoni,
- G. (2003). Patterns and perceptions of complementary/alternative medicine among paediatricians and patients' mothers: A review of the literature. *European Journal of Pediatrics*, 162(12), 820–827.
- <sup>562</sup> De Barra, M. (2017). Reporting bias inflates the reputation of medical treatments: A
- comparison of outcomes in clinical trials and online product reviews. Social Science &
   Medicine, 177, 248–255.
- <sup>565</sup> De Barra, M., Eriksson, K., & Strimling, P. (2014). How feedback biases give ineffective <sup>566</sup> medical treatments a good reputation. *Journal of Medical Internet Research*, 16(8),

567 e193.

- <sup>568</sup> Edgerton, R. B. (1992). Sick Societies: Challenging the Myth of Primitive Harmony. New
- 569 York: Free Press.
- Fabrega, H. (1997). Earliest phases in the evolution of sickness and healing. Medical An thropology Quarterly, 11(1), 26–55.
- Finlay, B. L., & Syal, S. (2014). The pain of altruism. Trends in cognitive sciences, 18(12),
  615–617.
- Gardner, A., & West, S. A. (2006). Demography, altruism, and the benefits of budding.
   Journal of Evolutionary Biology, 19(5), 1707–1716.
- Glenton, C. (2003). Chronic back pain sufferers—striving for the sick role. Social Science &
   Medicine, 57(11), 2243–2252.
- Godfray, H. C. J. (1991). Signalling of need by offspring to their parents. *Nature*, 352, 328–330.
- Grafen, A. (1990). Biological signals as handicaps. Journal of Theoretical Biology, 144 (4),
  517-546.
- Gun, R. T. (1990). The incidence and distribution of RSI in South Australia 1980-81 to 1986-87. The Medical Journal of Australia, 153(7), 376–380.
- Hamilton, W. (1964). The genetical evolution of social behaviour. I. Journal of Theoretical Biology, 7(1), 1–16.
- Hamilton, W. D. (1963). The evolution of altruistic behavior. The American Naturalist,
   97(896), 354–356.
- <sup>588</sup> Hardy, A. (2006). First do no harm. *EMBO reports*, 7(12), 1199–1199.
- Henrich, J., & Henrich, N. (2010). The evolution of cultural adaptations: Fijian food taboos
   protect against dangerous marine toxins. *Proceedings of the Royal Society B: Biological Sciences*, 277(1701), 3715–3724.
- <sup>592</sup> Hill, K., & Hurtado, A. M. (2009). Cooperative breeding in South American hunter– <sup>593</sup> gatherers. *Proceedings of the Royal Society B: Biological Sciences*, 276, 3863–3870.
- Jiménez, Á. V., Stubbersfield, J. M., & Tehrani, J. J. (2018). An experimental investigation
- <sup>595</sup> into the transmission of antivax attitudes using a fictional health controversy. *Social*
- 596 Science & Medicine, 215, 23–27.
- <sup>597</sup> Kaplan, H., Hill, K., Lancaster, J., & Hurtado, A. M. (2000). A theory of human life history
- evolution: Diet, intelligence, and longevity. Evolutionary Anthropology Issues News and

- $_{599}$  Reviews, 9(4), 156-185.
- Kleinman, A. (1988). The Illness Narratives: Suffering, Healing & the Human Condition.
   *New York, USA: Basic Books*, 18–30.
- Marks, N. F. (1996). Caregiving across the lifespan: National prevalence and predictors.
   *Family Relations*, 27–36.
- Maynard Smith, J. (1991). Honest signalling: The Philip Sidney game. Animal Behaviour, 42(6), 1034–1035.
- Maynard Smith, J., & Price, G. R. (1973). The logic of animal conflict. Nature, 246, 15–18.
- Miton, H., & Mercier, H. (2015). Cognitive obstacles to pro-vaccination beliefs. Trends in cognitive sciences, 19(11), 633–636.
- Miton, H., Claidière, N., & Mercier, H. (2015). Universal cognitive mechanisms explain the cultural success of bloodletting. *Evolution and Human Behavior*, 36(4), 303–312.
- Mittenberg, W., Patton, C., Canyock, E. M., & Condit, D. C. (2002). Base rates of malinger-
- ing and symptom exeggeration. Journal of Clinical and Experimental Neuropsychology,
   24(8), 1094–1102.
- <sup>614</sup> Nicholson, K., & Martelli, M. F. (2007). The effect of compensation status. In G. Young, A.
- <sup>615</sup> W. Kane, & K. Nicholson (Eds.), *Causality of Psychological Injury: Presenting Evidence* <sup>616</sup> *in Court* (pp. 411–426). New York: Springer.
- Norman, G. R., Tugwell, P., & Feightner, J. W. (1982). A comparison of resident performance on real and simulated patients. *Academic Medicine*, 57(9), 708–715.
- Nowak, M. A., & Sigmund, K. (1998). Evolution of indirect reciprocity by image scoring.
   *Nature*, 393(6685), 573.
- Parsons, T. (1951). Illness and the role of the physician: A sociological perspective. Ameri *can Journal of orthopsychiatry*, 21(3), 452–460. Wiley Online Library.
- Parsons, T., & Fox, R. (1952). Illness, therapy and the modern urban american family.
   Journal of Social Issues, 8(4), 31–44.
- Perkins, M., Howard, V. J., Wadley, V. G., Crowe, M., Safford, M. M., Haley, W. E., Howard,
   G., et al. (2012). Caregiving strain and all-cause mortality: Evidence from the regards
- study. The Journals of Gerontology Series B: Psychological Sciences and Social Sciences,
- $628 \quad 68(4), 504-512.$
- Raihani, N., & Bshary, R. (2011). Resolving the iterated prisoner's dilemma: Theory and
   reality. Journal of Evolutionary Biology, 24 (8), 1628–1639.
- Rohling, M. L., Binder, L. M., & Langhinrichsen-Rohling, J. (1995). Money matters: A
   meta-analytic review of the association between financial compensation and the experi-

- ence and treatment of chronic pain. Health Psychology, 14(6), 537.
- Rousset, F., & Billiard, S. (2000). A theoretical basis for measures of kin selection in sub divided populations: finite populations and localized dispersal. *Journal of Evolutionary Biology*, 13(5), 814–825.
- Schulz, R., & Beach, S. R. (1999). Caregiving as a risk factor for mortality: The caregiver
  health effects study. JAMA, 282(23), 2215–2219.
- Steinkopf, L. (2015). The signaling theory of symptoms an evolutionary explanation of the
  placebo effect. *Evolutionary Psychology*, 13(3), 1474704915600559.
- Steinkopf, L. (2016). An evolutionary perspective on pain communication. *Evolutionary Psychology*, 14(2), 1474704916653964.
- Steinkopf, L. (2017). The social situation of sickness: An evolutionary perspective on therapeutic encounters. *Evolutionary Psychological Science*, 3(3), 270–286.
- <sup>645</sup> Sugg, R. (2008). Corpse medicine: Mummies, cannibals, and vampires. *The Lancet*,
   <sup>646</sup> 371 (9630), 2078–2079.
- Sugiyama, L. S. (2004). Illness, injury, and disability among Shiwiar forager-horticulturalists:
   Implications of health-risk buffering for the evolution of human life history. American
   Journal of Physical Anthropology, 123(4), 371–389.
- Tanaka, M. M., Kendal, J. R., & Laland, K. N. (2009). From traditional medicine to
  witchcraft: Why medical treatments are not always efficacious. *PLoS One*, 4(4), e5192.
- Taylor, P. D., & Frank, S. A. (1996). How to Make a Kin Selection Model. Journal of
   Theoretical Biology, 180(1), 27–37.
- <sup>654</sup> Tiokhin, L. (2016). Do symptoms of illness serve signaling functions? (Hint: Yes). *The* <sup>655</sup> *Quarterly Review of Biology*, 91(2), 177–195.
- Trivers, R. L. (1971). The evolution of reciprocal altruism. The Quarterly Review of Biology, 46(1), 35-57.
- <sup>658</sup> Trivers, R. L. (1974). Parent-offspring conflict. *American Zoologist*, 249–264.
- <sup>659</sup> Trnka, S. (2007). Languages of labor: Negotiating the real and the relational in Indo-Fijian <sup>660</sup> women's expressions of physical pain. *Medical Anthropology Quarterly*, 21(4), 388–408.
- Vitaliano, P. P., Zhang, J., & Scanlan, J. M. (2003). Is caregiving hazardous to one's physical
  health? A meta-analysis. *Psychological Bulletin*, 129(6), 946.
- <sup>663</sup> Wessely, S. (2003). Malingering: Historical perspectives. In P. W. Halligan, C. Bass, &
- <sup>664</sup> D. A. Oakley (Eds.), *Malingering and illness deception* (pp. 31–41). Oxford: Oxford

- 665 University Press.
- <sup>666</sup> West, S. A., Pen, I., & Griffin, A. S. (2002). Cooperation and competition between relatives.
- 667 Science, 296(5565), 72-75.
- <sup>668</sup> Withey, A. (2013). Physick and the family: Health, medicine and care in Wales, 1600-1750.
- 669 Oxford: Oxford University Press.
- <sup>670</sup> Wootton, D. (2006). Bad medicine: Doctors doing harm since Hippocrates. Oxford: Oxford
- 671 University Press.
- Zahavi, A. (1975). Mate selection: A selection for a handicap. Journal of Theoretical Biology,
   53(1), 205–214.