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# LAY SUMMARY

When a person is diagnosed with cancer, they may wish to consider undergoing fertility preservation procedures. These procedures give patients a chance to have their own biological child after completing cancer treatment. However, research suggests that cancer patients who have a poor prognosis are less likely to be offered fertility preservation treatment. In this paper, we consider the ethical implications of offering (or not) fertility preservation to this patient group, including using their sperm or eggs to reproduce after their death. We conclude that fertility preservation treatments should be offered to all cancer patients who might benefit from it, and we outline the many ways that benefit from this treatment can be gained. The decision to withhold the offer of fertility preservation treatment should be made between the patient's clinician and their wider care team. They must be able to provide good reasons to explain why it was withheld.

# Reproduction in life and death: should cancer patients with a poor prognosis be offered fertility preservation interventions?

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- 4 Authors: Georgina L Jones<sup>1</sup>, Anne-Mairead Folan<sup>1</sup>, Bob Phillips<sup>2,3</sup>, Richard A Anderson<sup>4</sup>,

5 Jonathan Ives<sup>5</sup>.

- <sup>6</sup> <sup>1</sup>Department of Psychology, School of Humanities and Social Sciences, Leeds Beckett
- 7 University, Leeds, LS1 3HE, UK.
- 8 <sup>2</sup> Hull-York Medical School and Centre for Reviews and Dissemination, University of York,

9 York, UK

- 10 <sup>3</sup> Paediatric Oncology, Leeds Children's Hospital, Leeds, UK
- <sup>4</sup>MRC Centre for Reproductive Health, University of Edinburgh, Edinburgh, UK
- 12 <sup>5</sup>Centre for Ethics in Medicine, University of Bristol, Bristol, UK
- 13
- 14 To whom correspondence should be addressed: Professor Georgina Jones, Department of
- 15 Psychology, School of Humanities and Social Sciences, Leeds Beckett University, Portland
- 16 Building, City Campus, Leeds. Tel +44 113 8125106. Email g.l.jones@leedsbeckett.ac.uk

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# 19 ABSTRACT

20 In the context of a cancer diagnosis, fertility preservation interventions are used to mitigate the 21 potential impact of gonadotoxic cancer treatment upon fertility. They provide patients with 22 cancer the option to freeze their reproductive material to have their own biological child 23 following treatment. The evidence suggests some clinicians are less likely to have fertility 24 preservation discussions with patients who have an aggressive or metastatic cancer which has a poor prognosis. Although this is contrary to current policy recommendations, there is a lack 25 26 of guidance relating to offering fertility preservation in the context of a poor prognosis to 27 support clinicians. Controversy surrounds posthumous reproduction, and whether the wishes 28 of the cancer patient, when living and deceased should take precedence over others' wellbeing. 29 We consider the question of whether cancer patients with a poor prognosis should be offered 30 fertility preservation from an ethics perspective. We structure the paper around key arguments 31 to which multiple ethical principles might pertain, first establishing a central argument in 32 favour of offering fertility preservation based on respect for autonomy, before exploring 33 counterarguments. We conclude by proposing that a defeasible assumption should be adopted 34 in favour of offering fertility preservation to *all* cancer patients who might benefit from it. It is 35 important to recognise that patients could benefit from fertility preservation in many ways, and 36 these are not limited to having a parenting experience. The burden of proof rests on the clinician 37 in collaboration with their multi-disciplinary team, to show that there are good grounds for 38 withholding the offer.

39

#### 40 LAY SUMMARY

When a person is diagnosed with cancer, they may wish to consider undergoing fertility 41 42 preservation procedures. These procedures give patients a chance to have their own biological 43 child after completing cancer treatment. However, research suggests that cancer patients who 44 have a poor prognosis are less likely to be offered fertility preservation treatment. In this paper, 45 we consider the ethical implications of offering (or not) fertility preservation to this patient 46 group, including using their sperm or eggs to reproduce after their death. We conclude that 47 fertility preservation treatments should be offered to all cancer patients who might benefit from it, and we outline the many ways that benefit from this treatment can be gained. The decision 48 49 to withhold the offer of fertility preservation treatment should be made between the patient's

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# 71 INTRODUCTION

Cancer treatments (e.g., chemotherapy, radiotherapy and some surgery) can result in loss of fertility - denying cancer patients the opportunity to have their own biological child in the future. Loss of fertility does not affect all patients treated for cancer (ESHRE, 2020), but predicting who will be affected is challenging because cancer treatments variably affect reproductive function depending on the patient's age, cancer diagnosis, and specific treatment regimen.

Procedures are now available which, if undertaken before cancer treatment, may preserve fertility (such as egg, embryo, sperm and ovarian/testicular tissue cryopreservation). Given that loss of fertility is often reported by cancer patients to be one of the most distressing side-effects of cancer treatment (Peate et al., 2009), fertility preservation (FP) options can provide much needed hope at a time when they are coping with significant uncertainty.

Crucially, the value of FP procedures lies in their ability to preserve the opportunity to have, 83 84 and to then parent, genetically related children, and the best chances of success for these procedures are when they are offered prior to commencing cancer treatment. Consequently, at 85 86 cancer diagnosis, international guidelines recommend that cancer teams should discuss the 87 impact of cancer treatment on future fertility with the patient (National Institute of Health and 88 Care Excellence 2017; ESHRE, 2020; Loren et al., 2013; Lambertini et al., 2020; Yasmin et 89 al., 2018). Ensuring that patients are supported to make the right FP decision for them is 90 essential, because patients surviving cancer will live with the consequences of the choices made 91 about their fertility for the rest of their lives, so it is vital that they are supported to make the 92 right FP decision for them.

Some cancer patients will have a poor prognosis, perhaps the result of a cancer which responds poorly to treatment, or one with widespread metastasises to other parts of the body, and there is little chance of cure. In these circumstances, the requirement to discuss FP options seems less clear, as patients in this position are very unlikely to be able to use their stored material to have and raise children.

98 Instead, it may be possible to use posthumous assisted reproduction (PAR), whereby 99 cryopreserved eggs, embryos or sperm are used after the death of an individual for the purpose 100 of expanding a family (Lawson et al., 2016). The notion of posthumous reproduction was first proposed by Montegazza in 1866, who first discovered that sperm could be frozen and
suggested that women whose husbands may have died during war could benefit from this
discovery (Elliot, 2004).

104 In these circumstances, where posthumous reproduction is the only likely option, we consider 105 the question of whether a FP discussion should take place and whether cancer patients with a 106 very poor prognosis should be offered fertility preservation.

107 Current available guidance for clinical teams is unclear on this question. Whilst there appears 108 to be a consensus that cancer teams should discuss the impact of cancer treatment on future 109 fertility with the patient at diagnosis, some professional and regulatory bodies such as The 110 American Society of Clinical Oncology (Lee et al., 2006) and the American Society for 111 Reproductive Medicine (2005) state that this should also apply to those with a poor prognosis, 112 and discussion should take place irrespective of prognosis in female cancer patients of reproductive age (Munoz et al., 2016, Loren et al., 2013, Peccatori et al., 2013, Lambertini et 113 114 al., 2016). Others such as the NICE Clinical guideline [CG156] in the UK and ESHRE recommendations (2020) do not. However, the NICE guideline does state that when deciding 115 116 to offer FP to people diagnosed with cancer, cancer teams should take into account factors including diagnosis and prognosis, and the viability of stored or post-thawed material. This 117 118 distinction might reflect differences between publicly vs self-funded health systems rather than 119 different ethical commitments, but it nonetheless provides evidence of varied positions and 120 practices that warrant ethical scrutiny.

To provide that ethical scrutiny, we first consider what the *purpose* of FP is, which we then use as backdrop against which to consider various arguments for or against offering FP to late stage or prognostically poor cancer patients.

#### 124 What is the purpose and process of fertility preservation?

125 It is important, at the outset, to consider what the purpose of FP treatment in the context of a 126 cancer diagnosis is, because this allows us to clearly articulate the goods that purportedly flow 127 from it. Whilst our aim is not to undertake a simple consequentialist analysis that balances 128 goods against harms, it is nonetheless important to have a clear sense of how people benefit 129 from FP and why it is valuable to them. 130 It seems that there are two kinds of goods that can flow from FP. The first is the good of being 131 able to reproduce genetically, and the second is the ability to be a parent to those genetically 132 related children. The former is independent of the latter, by which we mean it is possible to 133 reproduce genetically without having the experience of parenting the resulting children. In 134 contrast, the latter is dependent on the former. One cannot have the good of parenting 135 genetically related children without first reproducing genetically.

We contend that the purpose of FP, in the context of cancer treatment, is to allow people to 136 experience the second good. The benefit derived from offering FP to patients undergoing 137 138 cancer treatment is that it leaves open a future where they could be a parent to their genetic 139 offspring. This seems clear for two reasons. First, if we were not concerned with preserving 140 the opportunity to parent genetic offspring specifically, we would not be concerned about 141 fertility preservation at all. We would simply reassure patients that notwithstanding the likely 142 loss of fertility, they will have the opportunity to parent non-genetically related children through gamete donation or adoption. Second, although it is possible to simply preserve 143 144 reproductive material so that it might be used at some point in the future by *someone*, which 145 would be enough to actuate the first good, the fact that people tend to want to preserve their 146 material for their own use, rather than to donate, suggests that they do not simply want to 147 reproduce, but that they are looking for a parenting experience. That said, we feel it is 148 reasonable to say that the primary aim of FP is to preserve a patient's ability to experience 149 being a parent to their genetically related child, as opposed to merely being a genetic progenitor 150 in the absence of a parenting experience. Thus this includes the possibility of using surrogacy 151 where, while another person carries the pregnancy, the original patient has both a genetically 152 related child and a parenting experience.

Of course, in the context of an adult patient with a partner, the aim of FP can also be to preserve the opportunity for that partner to have the experience of being a parent to children genetically related to the patient and the surviving partner. Consistent with what we have said above, we contend that the good derived from this is that of enabling the surviving partner to have a parenting relationship in a way that connects them to the deceased patient, rather than the good of simply having genetically reproduced (which could be achieved by donation to a stranger).

Having now established that the purpose of FP is not simply to facilitate genetic reproduction
 *qua* genetic reproduction, but rather to preserve the option of having a parenting experience

with one's genetic progeny for either oneself or one's partner, we will go on to considerarguments for and against routinely offering FP.

#### 163 **Respecting autonomy**

Respecting autonomy is a cornerstone of Western medical ethics, and this requires us to respect a person's right to make decisions for themselves, including about their medical care. Respect for autonomy does not mean we must do whatever people want, but that we take their wishes seriously and allow them to choose when a choice is available. In order to act autonomously, people need to be sufficiently informed about what their options are and the likely consequences of decision options - a fundamental prerequisite for ensuring informed consent.

Increasingly, patients (including cancer patients) expect to participate and be involved in their treatment decisions (Brietsameter, 2010; Siminoff and Thomson, 2010). Shared decisionmaking is a process whereby clinicians share information about treatment options, empowering the patient to actively decide based on their preferences (Elwyn et al., 2012) and this process is considered a quality benchmark for the delivery of dignified care (Department of Health, 2012; Coulter and Collins, 2011).

As such, in order to respect autonomy, a high-quality fertility discussion in oncology is critical to ensure that cancer patients are sufficiently informed about the potential impact of cancer upon fertility. This will support them to make autonomous, fully informed decisions that give them control over their reproductive future, before giving consent to any subsequent intervention.

181 Hudson et al. (2016) have noted that the need to respect patient's freedom of thought, intention and action while delivering healthcare, seems to suggest that everyone should be offered FP. 182 This seems plausible. An autonomy-based argument can clearly be made for having a FP 183 184 conversation with every patient or the patient's proxy decision maker (i.e. a parent) if they are not able to make a decision by virtue of their age, or a best interest decision made in accordance 185 with the Mental Capacity Act (2005) if they are an adult who is otherwise incapacitous. Not 186 to raise the issue of fertility loss, and inform patients of FP options, would be a *de facto* 187 188 deception by omission and this is an insult to autonomy.

189 Hudson et al. (2016) do sound word of caution about drawing this conclusion, however, asking 190 whether cancer patients with a poor prognosis are really able to make autonomous decisions? 191 For example, the extent to which cancer patients can actually make informed decisions 192 regarding post-humous reproduction PAR has been debated, as a result of the time pressure 193 needed to make a FP decision and emotional and cognitive factors which may inhibit the ability 194 to retain information (Lawson et al., 2015). In our view, whilst it is of course possible that some 195 patients in this position will not have the capacity to make a complex future looking decision 196 about their fertility, capacity is person and decision specific, and no blanket assumption can be 197 made.

198 That said, although there is a strong prima facie argument for respecting autonomy by giving 199 (capacitous) patients the option of FP, various studies suggest that doctors aren't providing all 200 patients with information about FP (Meyer and Farrell, 2015). Many healthcare professionals 201 report a decreased likelihood of initiating discussions related to FP with patients who have a 202 poor prognosis. Poor prognosis has been cited as a major barrier for talking about FP with 203 patients by 66.9% of oncologists (Zhang et al., 2020), with other studies citing that over 50% 204 of oncologists state that a patient's poor prognosis may dissuade them from discussing FP 205 (Sallem et al., 2018; Forman et al., 2009). Comparable findings have been reported in other 206 studies, with 41%-88% of clinicians citing poor prognosis as a factor that would either 207 influence FP discussions or lead them to not offer the option to patients (Chung et al., 2017; 208 Adams et al., 2013; Louwé et al., 2013; Collins et al., 2011). Similarly, a survey of paediatric 209 oncologists' attitudes and practices toward FP in adolescents reported that a patient having a 210 poor survival prognosis was one of three most likely reasons for the physician to not 211 recommend sperm banking (Köhler et al., 2011).

Quinn et al. (2009) reported that the majority of oncologists included in their research do not discuss FP with patients with a poor prognosis. One oncologist reported that they experienced discomfort at the thought of talking about "future babies" with a patient that is unlikely to be alive within months. The authors concluded that although guidelines suggest healthcare professionals should discuss FP with all patients, the majority appear to not follow these guidelines. Takeuchi et al. (2017) observed an internal conflict in clinicians regarding whether, and when, it was appropriate to discuss FP with patients with a poor prognosis.

This evidence about clinician practice tallies with studies reporting patient experience, with Jones et al. (2022) reporting patients saying they were not informed about or offered FP at 221 cancer diagnosis. One explanation for this may be that clinician preferences can be limited by 222 'implicit persuasion'. This is a process whereby clinicians subconsciously place greater 223 emphasis on the treatment options they consider to be better suited to the patient, a phenomenon 224 frequently observed in oncology settings (Englehardt et al., 2016). When an oncologist's goal 225 is to save lives and treat the cancer, they may see that as taking precedence and discussion 226 about FP a distraction that could delay treatment and thus worsen prognosis at the detriment to 227 the patient (Hudson et al., 2016). This may be a conscious omission (e.g. due to the pressure to 228 initiate cancer treatment), or an unconscious bias. The former could be either an unethical 229 insult to autonomy or an act of beneficent paternalism, whereas the latter would seem to be a 230 form of negligent or unprincipled omission.

231 Whilst some clinicians may take a conscious position against offering FP, or simply not think 232 to mention it, a number of studies have reported that many healthcare professionals take a 233 neutral stance or have a mixed responses to fertility issues in cancer patients with a poor 234 prognosis. Quinn et al. (2012)'s findings revealed that 45.2% of oncologists had a neutral 235 stance on the issue of patients with a poor prognosis pursuing FP. When questioned about 236 posthumous parenting, 16.2% of oncologists expressed their support of this, whilst the majority 237 (51.5%) did not have an opinion on the matter. Oncologists who were aware of the American 238 Society of Clinical Oncology guidelines (which highlight the importance of patient education 239 and acknowledge the need for oncologists to provide fertility-related information) were more 240 likely to discuss fertility issues with patients with a poor prognosis. Rosenberg et al. (2017) 241 found that 58% of oncologists believed that patients with a poor prognosis should not pursue FP. However, overall, this belief did not appear to dissuade the majority of oncologists from 242 discussing fertility issues with this patient group. When asked whether they discuss fertility 243 244 issues with this patient group, 84% of oncologists reported that they either always, or usually, 245 had these discussions (43% and 41%, respectively), whilst 15% rarely did and 1% never had 246 these discussions.

Some studies suggest that nurses may be more inclined to discuss FP with all patients, regardless of prognosis. In a study by Vadaparampil et al. (2007), paediatric oncology nurses were asked about discussing FP options with patients. The majority (68%) of nurses stated that a patient's poor prognosis would not affect the discussion being carried out. Just over one quarter (28%) of nurses stated that they would be less likely to discuss FP options with this group of patients, and 4% stated that a patient's poor prognosis would make them more likely 253 to discuss FP options. Comparable findings were observed by Krouwel et al. (2017), who 254 carried out a survey study with 421 oncology nurses to investigate their knowledge about FP 255 and possible barriers to discussions of this topic. When asked to rate their agreement with the 256 statement: "I would tend not to discuss FP with a patient because the patient has poor 257 prognosis", over half of nurses (55.2%) strongly disagreed with this statement. Over a quarter 258 (28.4%) of participants stated that they neither agreed nor disagreed with this statement, with 259 16.4% strongly agreeing with the statement. Similarly, in a study comparing nurses' views on 260 FP in patients with a poor prognosis between 2006 and 2005 (Clayton et al., 2008), regardless 261 of the survey year, the majority of nurses reported that a poor prognosis would not affect the 262 likelihood of FP discussions. The number of nurses reporting that FP discussions were more 263 likely for patients with a poor prognosis also increased from 2005 (5%) to 2006 (22%), in 264 addition to a slight decrease in the number of nurses reporting that they would be less likely to discuss FP with patients with a poor prognosis (this figure was 27% in 2006 and 28% in 2005). 265 However, King et al. (2008) reported that the odds of a patient surviving was a factor that 266 267 determined whether nurses discussed FP with their patients.

What we can take from this is that there is mixed practice, but whereas the clinicians who always offer FP will always be respecting autonomy, those who never or only sometimes do seem to be acting, at least *prima facie*, unethically, in light of the strong autonomy argument outlined above. The question, then, is whether these data evidence an ethical failure on the part of many clinicians, or whether this variation in practice should give us pause to ask whether there are valid reasons that override the *prima facie* autonomy argument. It is the latter possibility that we will now explore.

#### 275 Justice and Opportunity Cost

276 The argument from justice and opportunity cost is concerned with resource prioritisation, and might contend that FP, in cases where it will not lead to a parenting experience, is futile and 277 278 therefore not a good use of resource. Crystalising this more generally for a publicly funded 279 health system, where treatment prioritisation decisions will be based on cost-effectiveness 280 metrics, this is because the decision to fund FP where the prognosis is poor cannot be justified 281 at a societal level as a reasonable distribution of scare resources. It can also plausibly be argued 282 that having genetically related children, whilst very important to many people, is not a need but rather a desire (McTernan, 2015). Even without taking a position on that, however, it seems 283 284 that FP cannot achieve its aims unless it has a chance of resulting in a parenting experience -

which it does not if the prognosis is very poor, and the patient is unlikely to survive to be a parent. This argument would, then, suggest that FP should not be offered to patients where FP will not result in a meaningful parenting experience for either them or their partner – and so would rule out FP for children and single adults who are unlikely to survive to become parents.

This perhaps gives us reason to revisit our previous claims about the purpose of FP. If we are going to rule out FP for some patients on the basis of futility, we ought to be sure that there are not other reasons that might justify providing it, even when a parenting experience cannot be achieved.

#### 293 Hope and Imagined Futures

There are two linked arguments to consider here, one centred around FP providing hope for a future, and the other around an imagined future as a parent.

#### 296 Hope for a future

297 It is well evidenced that the experience of infertility can lead to distress which can influence the individual emotionally in the short-term, and also their sense of identity and expectations 298 299 for the future (Letherby, 1999). Franklin (2022) has suggested that IVF and its related 300 technologies are 'hope technologies', because they offer a seductive techno-solution to the 301 enigmatic problem of infertility, and in this context, clinicians become providers of hope. It 302 can be argued that for cancer patients with a poor prognosis, FP treatments, which will often 303 involve assisted reproduction techniques, are also hope technologies. Hope has also been 304 conceptualised and operationalised by Snyder (1995), who contends that its existence is 305 essential as a psychological coping strategy, and it could be argued that everything that can be 306 done to better support poor prognosis patients at this time should be done. However, cognitive 307 rules which govern the appropriateness of hope include such criteria as the goal being under 308 some control and at the mid-range in terms of probability (Averill et al., 2012).

When the prognosis is good, it is not contentious to assume that offering FP treatment is appropriate because it is nurturing realistic hope of future parenthood. In contrast, where patients have a poor prognosis, where even the probability of using the stored material is low (yet alone success rates from any subsequent IVF/surgery), it may be deemed that nurturing hope by offering procedures, in these circumstances, is considered inappropriate. Furthermore, the act of offering FP to a patient with a very poor prognosis might itself offer unrealistic hope of survival, and may be interpreted as saying that there is good chance of living to become a parent – because why offer it otherwise? Indeed, Vadaparampil et al. (2008) found in their qualitative study that many of the 24 paediatric oncologists described how just mentioning the need for FP was seen as a sign of hope for patients and families.

The offer of FP, then, essentially becomes an offer of hope for a future which is false hope. It may be that in some cases hope, false or not, can provide benefit, and so becomes a benevolent lie – but in other cases giving false hope will be harmful, leading to a loss of trust and increased trauma when the falsehood becomes apparent. The potential harm arising from false hope exposed might be amplified if the retrieval intervention was burdensome to the patient, and the wrong amplified if it was costly to the health service or patient (if self-funded).

As such, we feel that offering FP to provide false hope as a therapeutic intervention is highly problematic, akin to a lie. Such a benevolent lie might on occasion be justified but would be the exception rather than the norm. However, it may be possible for a patient to benefit from an imagined future as a parent, even in the knowledge that they will never become one.

329 There is one further important argument to make here. It could be argued that the speed of 330 medical progress is high; so that what is an incurable cancer today may be treatable tomorrow. 331 Furthermore, clinical judgement is fallible and patients who are believed to be terminal may 332 not be. This could lead to the argument that we should offer FP treatment to everyone because 333 a) the terminal diagnosis may be wrong, and b) the currently terminally ill patients may end up 334 surviving due to advances in medical treatments. In both cases, the surviving patients will lose 335 their ability to reproduce if not offered FP. Therefore, not offering FP to a patient due to their 336 poor prognosis may not be justified. Some patients may survive and deeply regret their lost 337 fertility, while those who do not survive and have their fertility preserved are no worse off.

This argument has some merit, but it is grounded in the assumption that it is preferable to provide a costly service to ensure that no-one might miss an opportunity to parent, and it is not clear that this is a proportionate or justifiable response, especially when the potential benefit of a live birth is not guaranteed. The possibility of diagnostic error, or the possibility that cure may be around the corner, is not, in our opinion, enough to offset entirely the problems we have outlined. Rather, the possibility and likelihood of both would (and we suggest usually would) be routinely factored into prognostication and therefore into the decision about whetherto offer FP.

#### 346 Imagined future as a parent

347 One reason sometimes cited for pursuing FP when diagnosed with a cancer with a poor 348 prognosis, is the desire for a 'genetic legacy' (Hudson et al., 2016). As mentioned above, FP 349 can result in the production of an offspring that can continue the cancer patient's bloodline 350 through the process of post-humous reproduction (Quinn et al., 2012). There is some evidence 351 to suggest that the public view PAR favourably (Barton et al., 2012; Hans and Dooley, 2014), 352 although PAR may be viewed negatively by oncologists when it comes to teenage and young 353 adult and paediatric cancer patients (Quinn and Vadaparampil, 2011) and a therefore a barrier 354 to a FP discussion.

355 The argument from imagined futures is different to an argument from genetic legacy. As 356 mentioned above, there is reason to be sceptical about a genetic legacy motivation purely because people exploring FP do not tend to do so with the aim of donating their material for 357 358 just anyone to reproduce with (which would satisfy the need for genetic legacy), but for either 359 their own use or the use of someone they already have a relationship with. The argument from 360 imagined futures, instead, is based on the good that might come from the dying cancer patient, 361 who knows they will never have a parenting experience, taking comfort in imagining their 362 future child. For those patients where future parenthood had always been part of their sense of 363 identity and long-term goals and, then the comfort gained from taking concrete steps that allow 364 them to vividly imagine the existence of their child, and a parent-child relationship, may in 365 itself be enough to justify offering FP even to patients who know and understand they will 366 certainly not survive to parent.

This imagining might also include imagining a future in which their partner and child are flourishing and getting comfort from each other, which leads to the argument from gifting.

#### 369 Gifting

Here, a cancer patient might see FP as a way to gift the opportunity to become a parent to the

partner who will survive them, and with whom they may have planned to have children. Thismay be linked to the argument from imagined futures, but it need not be. It may simply be an

act of generosity, where they wish to gift something to someone they love, even though they

will never see or experience the result – and there seems little to object to ethically in such anact.

376 Something that ought to be considered in light of both of these arguments however, is the 377 pressure that cancer patients may feel under to act as though they value reproduction, and 378 engage with FP simply to please others because they feel it is expected of them. We live in a 379 pronatalist society, where reproduction is the norm (Greil et al., 2011), and in which people are more often required to explain their decision not to have children than their decision to have 380 381 them. Indeed, the act of simply offering FP is enough to suggest that it is important to preserve fertility. In this context, where people may feel under pressure to 'gift' their reproductive tissue 382 or act as if the thought of having children brings them comfort, it is important that if and when 383 384 FP is raised, it is presented as a genuinely neutral choice.

385 So far, we have considered the argument from autonomy, which gives us good reason to offer 386 FP to everyone, followed by the argument from justice and opportunity cost that suggests it 387 would be justifiable to not offer FP to patients with poor prognosis because they would never experience the goods of being a parent. We then considered two arguments, the argument from 388 389 imagined futures (including hope) and the argument from gifting, which support the idea that 390 patients may nonetheless wish to consider FP for reasons other than having a parenting 391 experience themselves. These latter two argument do, however, appear to limit the scope of to 392 whom FP should be offered, to include cancer patients who have a partner to whom they wish 393 to give the gift of becoming a parent, or anyone who would be comforted by imagining a 394 relationship with their future children (even if they will not experience it). This would seem to 395 exclude, however, in all cases, child patients, who would not have a partner to gift to, and seem 396 very unlikely to be able to imagine a parental relationship and gain comfort from it.

This exclusion of children with poor prognosis from FP might be supported by other arguments, notably the need to protect them from undergoing interventions that will not benefit them, and which they might consent to undertake in order to please others, because they think they ought to do it (see above comment about pronatalism).

Essentially, there are good reasons to think that FP can and should be offered to patients with poor prognosis, but only when they have the capacity to benefit in some way. In the absence of that capacity to benefit, there is little justification for offering FP and indeed given the risk of the offer itself being pressuring there are prima facie reasons not to offer it. 405 So far, the arguments we have considered have been patient focussed, and it is worth briefly 406 considering some wider arguments that could impact on the summary conclusion outlined 407 above.

#### 408 The welfare of the subsequent child

Arguments based on the welfare of the child are sometimes made to either support or oppose
FP in this context. We do not find any of them convincing, and so will outline them only briefly
for completeness.

412 First, it might be argued that FP should not be offered to any patient with a poor prognosis 413 because any resultant child will be harmed by not having one of its parents. This argument can 414 be expressed in different ways. Hudson et al. (2016) and Quinn et al. (2012) note that there 415 could be economic and social harms to a child raised without one parent. Taking a different 416 line, Lawson et al. (2016) have suggested children conceived via posthumous reproduction can be compared to 'replacement children', a term used by researchers to describe children who 417 418 are born following the death of an older sibling and may therefore be placed under unreasonable 419 pressure to be a good replacement.

420 We would resist these arguments. First, they speculate about necessary harm to children raised 421 in one parent families, which is not corroborated by evidence. Studies have highlighted that 422 children raised to single mothers through donor insemination display fewer emotional and 423 behavioural difficulties than those raised by married couples who have used donor 424 insemination (Murray and Golombok, 2005), in addition to solo women reporting higher education levels, higher income professions, and equally or higher perceived social support 425 426 from friends and family in comparison to cohabiting women awaiting fertility treatment 427 (Pettersson et al., 2023). Second, it problematically assumes that the surviving partner would 428 never find another partner who would be a step-parent or that they will themselves be an unfit 429 parent. Third, it assumes that a life with only one parent is not only sub-optimal but is a life 430 not worth living – which seems very problematic and would be contested by many people so 431 raised. A longitudinal study exploring changing family dynamics in over 27,800 single parent 432 households reported no evidence of a negative impact from living in a one parent household 433 on children's wellbeing, with children reporting equally high, or higher, scores in various 434 measures of wellbeing compared to those who have always lived in a two parent household 435 (Rabindrakumar, 2018). Furthermore, if there is genuine concern about child welfare linked to

financial and home situations, then there should be financial and welfare thresholds for all people having children – not only where assistance is needed. We therefore dismiss this concern, and note that the ASRM guidelines state that concerns about the welfare of the offspring are not sufficient cause for denying FP (The Ethics Committee of the American Society for Reproductive Medicine, 2005).

441 Second, it might be argued that by not offering FP, possible children are harmed by not being 442 brought into existence. We raise this only to dismiss it quickly, as it does not seem feasible 443 that we could consider harm being affected to those who are never brought into existence, as 444 there is no subject that could be harmed.

#### 445 The welfare of the partner

Tremellen and Savulescu (2015) note that, in the context of posthumous reproduction, there could be implications for the welfare and quality of life of the partner, specifically in terms of raising a child in grief. Hudson et al. (2016) also consider this, noting that:

449 "…research is needed to explore the long-term impact on widowed partners and their offspring.
450 Partners in this precarious position should receive counselling and support during this
451 decision-making process."

There are also the general and ubiquitous challenges, both financial and emotional issues of single parenting. One immediate response is to note that a surviving partner is under no obligation to use the preserved material. But it is plausible to think that they may feel under pressure to use it *because it is there*. There may also be a financial cost of continuing storage. That said, this does not, in our view provide reason to not offer FP at all, but rather (as noted by Hudson et al., 2016) to ensure that a surviving partner is appropriately supported, and enters into, or chooses not to enter into, a posthumous parenting project autonomously.

### 459 **Concluding argument**

In the discussion above we have considered a range of arguments both for and against routinely offering fertility preservation to patients with poor prognosis. It is clear that there is a strong *pro tanto* argument, grounded in respect for autonomy, that supports routinely offering to all patients. We have, however, also shown that there are good reasons to sometimes be cautious 464 about making this offer, not least because there are some situations in which fertility 465 preservation would be futile and as such cannot be considered a good use of resource.

The way forward is not absolutely clear; however, we propose that we should adopt a defeasible assumption in favour of offering fertility preservation to all patients who might benefit from it, with the burden of proof on the clinician to show that there are good grounds for withholding the offer. Given that the main argument in favour of routinely offering FP is found in a patient's autonomous choice to benefit from the intervention, the most appropriate grounds for withholding the offer (outside of lack of resource) is if a patient cannot benefit from the intervention.

473 We must, however, recognise that there are many ways a patient could benefit from fertility 474 preservation, and these are not limited to having a parenting experience. Becoming a parent is 475 arguably the primary goal of fertility preservation, and is one clear benefit, but a patient may 476 also benefit from FP therapeutically in the form of gaining significant comfort from an 477 imagined future, or knowing they have gifted something precious to their partner. That said, the criteria for withholding the offer would hold if and only if FP could not achieve the benefit 478 479 that is sought. This would make FP a futile treatment, and therefore one which there is no 480 obligation to offer.

The challenge here, of course, is that a decision about futility that rests on a clinician's understanding of the patient's ability to benefit assumes that a clinician is able to correctly identify all the benefits that a patient might seek to achieve (requiring an intimate and accurate knowledge of the patient and of their partner where present), and accurately predict that these benefits cannot, or are very unlikely to, be achieved. The former would normally be difficult to do without having had a specific conversation with the patient about FP and the reasons why they might want it, and the latter would require great deal of prognostic certainty.

For this reason, we suggest that the presumption in favour of having a conversation with all patients, including those with very poor prognosis is strong, and can and should only be defeated when there is clear evidence, agreed by a multidisciplinary team, that no benefit is possible. In practice, this might describe a relatively rare situation. One obvious example might be a young child with a negligible to zero chance of survival, who lacks the capacity to benefit from an imagined future, and who is not in a position to want to make a gift to a partner. This describes a patient who cannot themselves benefit from FP, and therefore should not be offered it. This would hold true regardless of whether the child's parents wanted FP, because we assume - we think reasonably - that the justification for fertility treatment lies in the benefit to the patient and not to others, and a child in this position should not be used as a means to achieve a good for their parents. (Note that in the gifting argument FP is justified by the benefit to the patient of giving the gift of parenting to their surviving partner, not by the benefit that may accrue to the surviving partner - although they will certainly benefit as well).

Given that we are making an argument in favour of a strong presumption of offering FP, one 501 502 might ask why we do not go so far as to make an argument for it being obligatory. Such an 503 argument might be grounded both in respect for autonomy and a legitimate concern that 504 allowing clinicians any discretion in whether to offer the treatment opens up the possibility of 505 inconsistency and a postcode lottery, where access to treatment depends on the particular views 506 of the clinician in charge. The best way to combat this would be to ensure that every patient, 507 regardless of prognosis, is given equal access. The reason we do not adopt this position is 508 simply that we are more concerned with equity then equality. Nedha (2011, as cited in Paul, 509 2019) defines equality as treating each person the same, regardless of needs and requirements 510 (e.g., in this scenario, providing all patients with the option of FP, regardless of prognosis), 511 whereas equity can be described as treating each individual fairly depending on their needs. 512 Removing clinical discretion and making the offer of FP mandatory in all cases, would ensure 513 every patient is treated equally at a cost of the harms that can follow when the treatment offers 514 false hope, leads to a foreseeable waste of resource, or risks a vulnerable patient feeling 515 pressured into undertaking procedures from which they cannot benefit.

516 Our position does potentially place a great deal of power in the hands of the clinician as 517 gatekeeper, but this is mitigated by our requirement that the offer of treatment is a defeasible 518 presumption with the requirement of the MDT agreement if it is withheld. Placing the burden 519 of proof on the clinical team to show, and agree, that there is no possible benefit to the patient 520 protects the autonomous patient's right to make their own choices in very large part, whilst 521 permitting discretion in a few cases where it might be necessary and correct. In adopting this 522 position, we concur with Hudson et al. (2016) regarding the need to provide cancer and fertility 523 healthcare professionals with appropriate resources and training packages for addressing the 524 ethical and decision-making implications that arise in these scenarios, and to ensure that 525 informed consent processes are high quality and robust. For those patients who wish to proceed, 526 ongoing care should include appropriate counselling in relation to the possibility, or even

- 527 probability, of posthumous reproduction. It must be assumed that most patients in this position
- 528 will not survive, and their preferences for using their frozen gametes or embryos must be
- 529 clearly documented in line with relevant legislation and local regulations.
- 530 In outlining our arguments, we try to find an appropriate balance between ensuring equity of
- 531 opportunity to beneficial treatment, whilst allowing for the fact that in some cases FP will be a
- 532 waste of resource for no patient benefit.

# 533 **Declaration of interest**

GLJ, AMJ, BP and RAA declare that there is no conflict of interest that could be perceived as prejudicing the impartiality of the opinions reported. JI is member of the NICE Highly Specialised Technology Evaluation Panel. He has not been involved in any NICE discussions involving fertility or fertility preservation, and the views expressed and endorsed here are his and do not represent NICE.

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# 544 Author contribution statement

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546 GLJ conceived the manuscript, preparing the initial draft. GLJ, AMF and JI critically 547 developed this further, producing the first version, with additional contributions from BP and 548 RAA. All authors edited, reviewed and approved the final manuscript.

549

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