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Predicting Success in the Embryology Lab: The Use of Algorithmic Technologies in Knowledge Production

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Abstract

This article analyzes local practices in embryo assessment in light of the increased use of time-lapse (TL) imaging in fertility treatment. The data produced by TL technologies are expected to help professionals pick the best embryo for implantation. The emergence of TL has been characterized by promissory discourses of deeper embryo knowledge and expanded selection standardization, despite professionals having no conclusive clinical trial evidence that TL improves pregnancy rates. Our research explores how TL tools are used and conceptualized by clinic staff. We pay special attention to standardization efforts and knowledge-creation facilitated through TL and its incorporated algorithms. Using ethnographic data from 5 UK clinical sites, we argue that knowledge generated through TL is contingent upon complex human machine interactions that produce local uncertainties. Thus, algorithms do not simply add medical knowledge. Rather, they rearrange professional practice and expertise. Firstly, we show how TL changes lab routines and training needs. Secondly, we show that the human input TL requires renders the algorithm itself an uncertain and situated practice. This, in turn, raises professional questions about the algorithm's authority in embryo selection. The article demonstrates how, due to their situated nature, medical algorithmic technologies co-exist in tension with knowledge standardization.

Keywords: algorithm, embryology, embryo imaging, IVF, laboratory, standardization

Introduction: Embryo assessment, Algorithms and Knowledge Standardization

Time-lapse (TL) imaging technologies were developed in the midst of a research-intensive period in the field of assisted reproductive technologies (ART). Scientific developments in the area of embryo diagnostics are a result of stagnating success rates in fertility clinics – rates that have only improved slightly since the 1980s (Gleicher et al. 2019).

Although many seemingly ‘good-quality’ embryos are transferred every year to patients, ^{about} ~~roughly~~ only 30% of them result in a live birth, with minor variations by age ~~group~~, country and clinic (Kupka et al. 2014). TL was designed to mitigate uncertainties surrounding scientific knowledge of embryo quality and implantation potential (Adjuk and Zernicka-Goetz 2013; Montag et al. 2013; Schoolcraft and Meseguer 2017). TL’s ability to facilitate closer monitoring of embryos is supposed to result, in theory, in a more efficient selection process.

Although the current evidence on TL’s ability to improve pregnancy rates is not conclusive, clinic laboratories across the UK have incorporated imaging techniques in their work. **Our research shows how TL imaging and algorithmic practices produce local outcomes as a result of complex human-technology interactions. Thus, the initial promissory discourses of algorithmic streamlined knowledge are transformed through lab practice. The technology has required professionals to rearrange lab routines, while also navigating the demands of the algorithmic embryo selection process. We highlight how algorithmic knowledge is not simply added through the introduction of a new technology. The knowledge is rather co-produced alongside the practices that the introduction of TL disrupts. Consequently, lab professionals navigate algorithmic choices they perceive as subjective, thus questioning the role of TL as an authoritative knowledge source.**

Knowledge production and standardization in medicine have been topics of great interest for STS scholars (Cambrosio et al. 2006; Knaapen 2014; Moreira 2007; Timmermans and Epstein 2010). However, the ways in which standards are adopted or rejected in laboratory

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3 work (Doing 2004, 2008; Knorr-Cetina, 1995, 1998; Latour 1983) remain underexplored.
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5 Additionally, medical knowledge is increasingly reliant on algorithmic technologies, thus
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7 complicating existing relationship between knowledge-creation and laboratory practice. As
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10 new algorithmic technologies are introduced in biomedical contexts, it is also vital to probe the
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12 intersections between laboratory work, standardization and the messy heterogenous ways in
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14 which algorithms operate (Liu 2021; Ziewitz 2015). This article offers insight on these
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16 intersections by contributing both to emerging STS studies of the effects of algorithms and to
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18 the literature on medical knowledge production more widely.
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22 In a review of studies on standardization, Timmermans and Epstein (2010, 69) call for
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24 a “careful empirical analysis of the specific and unintended consequences of different sorts of
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26 standards operating in distinct social domains.” Although the rise of knowledge standardization
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28 can be observed in many fields, healthcare, in particular, has been undergoing a large
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30 standardization movement driven by the adoption of evidence-based-medicine (EBM)
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32 protocols during the past three decades. This has resulted in a focus on practice guidelines,
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34 increased standardization of outcome measures, as well as numerous meta-analyses medical
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36 literatures (Greenhalgh et al. 2008; Knaapen 2014; Moreira 2007; Timmermans & Berg 2003).
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38 The drive towards standardization, however, has also revealed the limitations of EBM. Clinical
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40 decision-making is still a complex process where professionals often make decisions based on
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42 local knowledge and experience (Berg, 1999; Greenhalgh et al. 2008). As STS scholars grapple
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44 with the social implications of algorithms (Crawford 2015; Lee and Harvey 2020), we offer a
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46 view of algorithmic lab technologies as situated and disruptive, suggesting that knowledge-
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48 creation through algorithms is a local process-in-the-making rather than a straightforward
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50 achievement through the introduction of a technology alone. More widely, we also suggest that
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52 scholars pay attention to the ways in which algorithmic technologies rearrange scientific
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54 practice.
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Embryo Knowledge and TL

Knowledge of human embryos remained limited until the late 1970s when ART began to be incorporated into medical practice (Chapko et al. 1995). Embryologists now know that optimal conditions can easily be disturbed when embryos are outside incubators. This creates tensions between the need to observe their development and the need to preserve them for successful implantation. Over time, embryologists developed a system where embryos are taken out of incubators for microscope observation only at specific points during their development, keeping disruption at a minimum. Before the introduction of TL, morphology (or how an embryo ‘looks’) was considered the best indicator of pregnancy potential (Holte et al. 2007). Although some studies indicate a correlation between ‘good-looking’ embryos and pregnancy rates, there are exceptions to this rule (Meseguer et al. 2012). Morphological assessment is a practice that continues due to tradition rather than robust evidence (Holte et al. 2007). It is sometimes categorized as ‘subjective’ due to inconsistencies between embryologist observations (Bendus et al. 2006). Social scientists (Helosvuori 2019) note that embryo assessment is achieved through the combination of several factors, including lab practices and professional expertise.

The introduction of TL is intended to mitigate knowledge uncertainty about embryo potential (Kaser and Racowsky 2014). TL’s non-invasiveness coupled with the routinization of live-cell imaging (DiCaglio, 2017; Landecker, 2012) have contributed to growing professional interest in it. Moreover, TL can process large amounts of embryo development data. There is a growing literature in embryology that uses these data. Although few definitive conclusions have been drawn, the promise of further standardization in embryo selection has gained traction in embryology (Lundin and Park, 2020).

In labs, TL use has introduced an additional criteria in embryo assessment: the timing of development events (e.g., nucleation, cell divisions). This facilitates a ‘morphokinetic’

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3 assessment of embryos which was associated with greater embryo viability in some exploratory
4 studies (Meseguer et al. 2012). TL also helps professionals detect abnormal events that can
5 occur in-between standard daily microscope observations (Freour et al. 2012; Wong et al.
6
7 2013). Certain abnormal embryo ‘behaviors’ (e.g., direct cleavage, where a cell divides into
8 three very quickly) are associated with lower implantation potential (Liu et al. 2014; Rubio et
9 al. 2012). The continuous embryo monitoring that TL provides is facilitated by incubator
10 cameras that take pictures every 5 to 20 minutes, resulting in a detailed development video. TL
11 software, however, is not yet able to detect developmental events automatically/through the
12 use of AI. Rather, embryologists are required to annotate these and recorded information in the
13 software so it can be processed by TL algorithms. Annotation involves embryologists watching
14 each embryo video very closely and marking the exact timing of embryo developmental events.
15
16 **When annotating, embryologists register these developmental events with a time-stamp in the**
17 **software. As such, they have to indicate exactly when the embryo reaches a particular stage,**
18 **for example, the appearance of the nucleus, the nucleus fading, ‘cleavage events’ (cell**
19 **divisions) and various embryo expansion stages (e.g., morula, blastulation). The later stages of**
20 **development are particularly hard to pin down precisely, according to embryologists. This is**
21 **because cells may often appear fuzzy or overlapping.**

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Within social studies of IVF, embryos themselves have been a contentious object in scientific research and practice (Ehrich et al. 2007; Parry 2006; Scott et al. 2012; Svendsen and Koch 2008; Van de Wiel 2018, 2019). Embryos are a locus of uncertainty in medical knowledge (Parry 2006; Scott et al. 2012) in addition to being entangled with moral debates regarding the beginning of life and the ethics of disposal (Ehrich et al. 2007; Svendsen and Koch 2008). In this article, we focus on embryos’ implantation potential and how this is assessed scientifically with the help of TL tools.

Working with TL

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6 Firstly, TL technologies consist of an incubator with cameras (optical microscope)
7 incorporated into its chambers (where the embryos are stored). One exception to this is the
8 PrimoVision brand which consists of a camera that can be attached to petri dishes in a standard
9 incubator. However, we found most labs prefer the cameras to be incorporated. A particular
10 brand ~~that~~ preferred by UK professionals is ~~Embryoscope~~. ^{Embryoscope is}
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16 Secondly, TL technologies
17 incorporate a software that allows the viewing of embryo images/videos on computers. The
18 software also incorporates the embryo selection algorithm that draws on the staff data input.
19 Although algorithms can differ and are customizable, their common purpose is to provide an
20 embryo grade. This can be used in conjunction with morphological assessment to determine
21 embryo implantation potential.
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29 Selection algorithms may differ slightly across clinics. One of the early TL systems
30 named Eeva was marketed as an AI-powered algorithm predicting which embryos are unlikely
31 to become a viable blastocyst (Kaser and Racowsky 2014). However, studies increasingly show
32 that universal selection algorithms are unlikely to work, as clinic populations vary. As such,
33 in-house personalized algorithms are preferred (Barrie et al. 2017; Fischer 2015). However,
34 developing a custom algorithm requires large data sets that not all clinics have yet. The
35 Embryoscope TL machines observed were often used in conjunction with a patented algorithm
36 package named KIDScore (Known Implantation Data score) used to: 1) deselect embryos that
37 behave abnormally, 2) predict likelihood of implantation on day 3 and on day 5 and 3) enable
38 clinics to develop their own algorithm following the collection of sufficient data on their
39 patients outcomes. Clinics observed ^d make use of KIDScore (although in different ways), while
40 also ~~working towards~~ building their own custom algorithm. If clinics use a different TL system
41 (such as PrimoVision or Eeva), a different annotation system and algorithm ^{are also} is used. However,
42 these systems are significantly less popular.
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Data and Methods

The data included in our analysis are part of a project studying the impact of imaging technologies in IVF. These include relevant medical guidelines and policy documents, lab observations and interviews with professional staff involved who have used TL. Professional observation and interview data were collected between June 2017 and March 2019. Detailed ethnographic observations were carried out by the authors in 5 England NHS sites (named here A, B, C, D and E) where fertility treatment is provided. We observed lab routines and shadowed embryologists at each site for a minimum of 3 working days. We paid close attention to the use of TL in the lab, the annotation and selection process and professional engagement with selection algorithms and information generated through TL technologies. The observations amounted to a total of 230 hours. Firstly, clinics were selected based on daily lab use of TL. Selection was also based on their availability and willingness to participate in the study. All 5 clinics agreed to participate and staff were informed in advance about study procedures, with all those observed signing a consent form prior to the start of our research. The study received university ethics approval as well as ethics clearance from the NHS and each clinic site. Following observations, professionals were approached by the authors regarding interviews. As with observations, interview participation was voluntary and involved the signing of an additional consent form. We conducted a total of 25 interviews. A small number of interviewees (e.g., clinic directors, nurses) were not lab staff, but had relevant TL knowledge or had talked to patients about its use. The majority of those interviewed and observed are, however, embryologists. The interviews lasted between 45 and 90 minutes, were audio recorded and then professionally transcribed. Our questions focused on participants' lab and work experiences, the challenges and benefits of using TL, and the technology's place in IVF treatment.

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3 TL has been consistently marketed on the basis that it can lead to improved rates of
4 pregnancy. This has been a definite factor into labs adopting it, but staff are also aware the
5 technology might not live up to its promise. Although clinics in our sample expressed that
6 technological hype and competitiveness in the IVF sector contributed to their adoption of TL,
7 they also stressed that they do not heavily market TL to patients and they do not charge extra
8 for it to be included in individuals' treatment. For this reason, commercialization issues did not
9 feature prominently in our data. Such issues have been explored in previous work (Van de Wiel
10 2018, 2019) and it is beyond the scope of this article to deal with them. In the analysis below,
11 we focus on how the technology has been rolled out locally in UK labs and how knowledge
12 creation is negotiated in practice.

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14 We analyzed the data using grounded theory principles (Glaser and Strauss 2017). ~~First,~~
15 we started with a set of initial codes based on the TL literature, ~~but we developed additional~~ ^{then}
16 ~~ones and~~ refined codes and grouping categories as the research progressed. The authors
17 constantly compared notes and observations that emerged from different research sites. The
18 situated practices that emerged from the data reveal that the use of TL is contingent upon
19 specific local procedures that problematize the TL standardization narrative. In the next section
20 we discuss the 'locality' of TL practices and uncertainty as they relate to annotation, the
21 algorithm and TL score use, as well as the sharing of TL embryo images with patients.

22 **Annotating Embryos: The Creation of New Lab Routines**

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24 The manual annotation process is a necessary precursor of TL algorithm output. One
25 resulting critique of TL is the increased need for professional consensus on how to annotate.
26 Annotating embryos is especially time-consuming. Thus, lab routines need to adjust for this
27 additional work created by the introduction of TL. The length of time required to annotate
28 varies depending on the quality of the embryo and professional experience. It also depends on
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3 how many embryos a patient has. For example, it is common for a patient to have 5-10 embryos
4 developing in the lab and sometimes more. For confident embryologists, the process can be
5 quicker. However, our observations revealed that consulting with other lab staff to reach
6 consensus on difficult annotations is part of the process and needed at least occasionally. We
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12 observed different annotation routines in each clinic, with each having to rearrange their
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15 practices to accommodate TL use.

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17 Firstly, clinics can make different choices regarding which embryos to annotate fully,
18 from fertilization to day 5. Embryologists at the biggest clinic (D) in our sample decided that
19 annotating all embryos would create unmanageable workloads, as annotating all could take
20 several hours daily for at least two staff. The embryos that are not annotated are usually those
21 that die early and are ~~thus~~ discarded. In some cases, only the ones that are good candidates for
22 transfer are annotated fully. In such cases, the goal is to have as much information about these
23 as possible, rather than collect data on all embryos. The lab director of a smaller clinic (B) and
24 TL use advocate, however, stressed that, for her, it is important to annotate all embryos in order
25 to take advantage of all data they provide. Consequently, she encourages staff to annotate all,
26 time permitting. This is possible when clinics have a manageable volume of patients. In clinic
27 B, we observed staff using time outside egg collection and transfer windows to catch up on
28 annotations. In all labs, we observed a preference for annotating either early in the morning
29 before egg collection procedures or later in the afternoon after patient appointments.

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47 In most clinics, annotation training is still ongoing. Some sites had a couple of staff
48 specifically tasked with embryo annotations. Others, however, annotate more widely and
49 conduct regular in-lab quality control exercises. To maximize TL benefits, annotation
50 consensus is needed. Nonetheless, during interviews, embryologists repeatedly emphasized
51 that some stages of embryo development might be harder to identify, thus leading to
52 'subjective' opinions on annotations. Depending on the level of TL integration, we observed
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3 more streamlined annotation consensus procedures in three of our clinics (B, C, and D).

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5 However, integration often came after an arduous training process. Importantly, embryologists
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7 talked about the changing scientific consensus:

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10 Well, it completely changed how you were as an embryologist. It was so ... And I, we got it when I
11 was mid-training so I'd gone from one way of doing it to oh no, now you need to learn it a completely
12 new way. *And the annotation is constantly changing. There's new things that we have to learn how to*
13 *annotate, definitions are changing, the consensus is changing all the time.* And so I do remember being
14 very, very late in the lab quite a lot trying to get my head round how to annotate and what to annotate
15 and yeah, it being quite difficult. And there's still people that struggle with it now. You know, that
16 definitely people find it really difficult. And also to see what the point of it is. You know, we annotate
17 about over 40 things on one embryo and we use a handful of them. So it has definitely increased
18 workload. And yes, my experience it was at the beginning very frustrating because it was a lot of, a lot
19 more work for what, for what benefit. And then I think that's what started me off on the well, there has
20 to be a reason why we're doing this. (Lab director, Clinic B)

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24 The lab director establishes the connection between TL-led changes and the uncertainty
25 surrounding annotation standardization. Various staff talked about the tediousness of keeping
26 up to date with medical literature developments. To a certain extent, TL has introduced another
27 learning curve in the lab, especially for those who were training when the machines became
28 popular. There is an optimistic caution in the professional community that TL integration will
29 deepen embryo knowledge. However, connecting this knowledge to lab practices requires
30 additional professional engagement with the scientific literature.

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40 We found that staff on the ground have to confront many questions regarding
41 consistency and quality control in TL practices. An embryologist at Clinic A, a clinic that has
42 not yet fully integrated TL explained:

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48 So at the moment there's only one or two people are annotating all of the embryos that are put into the
49 Embryoscope because they have been trained and they, their annotations have been compared to make
50 sure that they're similar or the same. So at the moment we are trying to train everyone to do be able to
51 do annotations but it's difficult to have, you know, a very cohesive, a very ... [interviewer: consistent?]
52 Yeah, consistent, that's the word I'm looking for. Consistent annotation. For things like cell divisions
53 it's fairly simple because you can see when it's divided or not but things like time to blastulation or
54 time to the start of, start of blastulation, so as soon as you can see a cavity appearing that's a little bit
55 subjective. Even with one operator it can vary but, or between embryos you can, your annotation time
56 might vary slightly so in that sense it does increase the workload slightly having the Embryoscope in
57 there. (Embryologist, Clinic A)

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3 With TL, labs have to create infrastructures for annotation quality control. As highlighted
4 above, some aspects are perceived as more subjective and thus, in need of standardization
5 (Timmermans and Epstein 2010). Objectivity, defined by staff as consistency in annotation,
6 was seen as an important pursuit meant to facilitate optimal use of TL. We observed staff
7 completing quality control exercises on a couple of occasions. Junior embryologists questioned
8 their annotations more, indicating that, to a certain degree, such skills are picked up through
9 repeated practice only. When asked what happens with inconsistent annotations, most staff said
10 that these are discussed with the person in charge of the exercise, in a process where they assess
11 why one person's annotations deviated significantly from expectations set by senior staff. Labs
12 also used a UK-specific external quality control exercise operated through the National
13 External Quality Assurance Service (NEQAS). TL videos are a relatively new feature for this
14 service. Most staff said that the quality of the TL videos provided by NEQAS was not as good
15 as the labs' own. The NEQAS exercise was usually managed in a similar way to the internal
16 exercise, where one senior member of staff was in charge of discussing inconsistencies.
17 Although such quality control exercises exist, staff stressed uncertainty in light of the need to
18 introduce new lab practices and rearrange how quality control for annotation is accomplished.
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42 Algorithms-in-the-making: TL and the Complex Human-Technology Interplay

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44 Our data reveal that TL is not a straightforward technological solution to standardizing
45 and mainstreaming embryo knowledge. In this section, we suggest that TL and algorithmic lab
46 technologies more broadly require careful unpacking, given their need to be activated through
47 professional input. Lab engagement with TL algorithmic platforms differs and is very much
48 dependent on lab expertise and willingness work on adapting software to lab practice. The use
49 of TL has exposed the need for additional expertise in the area of biostatistics – expertise that
50 is not typically built into fertility care. Nonetheless, labs across the UK that have adopted TL
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3 tools and have embraced the learning process required, although with different degrees of
4 enthusiasm. For example, in our sample, we found that at least two labs had overall reservations
5 regarding the benefits of using TL. Uncertainty regarding optimal use of algorithms featured
6 prominently in our discussions with embryologists. Staff were highly aware that, although the
7 technology holds promise, significant input was needed from them in order for the algorithm
8 to function at its full potential. This included annotation as well as setting up algorithm
9 parameters and embryo score outputs.

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11 In the initial stages of adopting an Embryoscope TL machine (used by all labs
12 observed), staff need to set up the KIDscore algorithm offered as an option for an extra cost.
13 The usage of the KIDscore package was seen by most as a practice in need of adapting to their
14 own clinic's needs. Algorithms developed outside of the clinic were often viewed with
15 suspicion:

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17 What I think might happen is that the undisturbed culture will maybe help. But then again I don't know,
18 I just don't know if we're using KIDscore to its full capacity for it to actually make a difference. And I
19 don't think anybody knows enough about KIDscore and enough about the algorithms of embryos to
20 actually say yeah, this is what you need to select the best embryo. I don't really trust it that much, that
21 algorithm. I do it because it's like we have to do it and whatever but it's very rare that we actually get
22 a higher KIDscore on what we would have thought was a lower quality embryo so usually they kind of
23 match up so I don't know, I don't know how much. And it's also very subjective, KIDscore. You know,
24 you're taking about when I think it's expanded and like you or someone else think it's expanded and it
25 can be completely different. So I think it's subjective too. So I don't think it's like ... I think again the
26 undisturbed culture and the idea of being able to look at it and you know, and you can see reverse
27 cleavage and stuff, that's quite interesting. But I don't know if it makes a big difference. (Senior
28 embryologist, Clinic B)

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30 The algorithmic black box creates knowledge uncertainty for lab staff that are not directly
31 involved with its creation or adaptation to their own clinical practice. This uncertainty was
32 perceived by participants as 'subjective' knowledge which they contrasted to an objectivity
33 ideal (or standard) that TL was meant to achieve. The need for extended human input into TL
34 was perceived as a source of subjectivity, thus problematizing the promise of TL as a
35 technology that could ensure a more seamless embryo selection process. Labs with research-
36 active embryologists who could coordinate the use of algorithm data were more confident with

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3 using TL. However, such practices create new skillset needs for some embryology labs. Despite
4 the promise of TL, its successful implementation on the ground depends on new expertise. As
5 the respondent above stresses, the annotation input in algorithms can still be categorized as a
6 ‘subjective’ endeavor as it requires embryologist consensus on visual data (e.g., the start of
7 embryo expansion) and is not automated. Thus, the management of uncertainty in relation to
8 TL use includes many variables, from the systematization of annotation procedures to the
9 setting of algorithm parameters. The constant need for human input into the machine was seen
10 as a main source of this uncertainty, thus rendering the technology somewhat incomplete in the
11 eyes of our participants.
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24 Uncertainty in local practice and technology-staff interactions was managed to some
25 extent by embryologists with TL research expertise. Two labs had at least one member of staff
26 with significant knowledge of TL algorithms. Nonetheless, the algorithm options seemed
27 daunting for most. For example, if a lab decides to build its own algorithm, there are a multitude
28 of directions to take with the annotations included, the weight given to different variables and
29 the inclusion of different patient conditions. This coupled with the constant need refine the
30 algorithm through the collection of new data. In small clinics, staff were weary of the long
31 transition to a robust algorithm, all the while knowing that it might not lead to increased
32 pregnancy rates. The uninterrupted incubation aspect of TL and the images generated through
33 it seemed to be more tangible benefits when compared to the uncertainty of algorithms and
34 their outputs.
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49 An illustration of the choices and human work involved into building TL algorithms
50 can be seen below. During lab observations, an embryologist from clinic C explained their
51 annotation and algorithm-building process:
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57 The senior embryologist says that Embryoscope has a variety of options for grading embryos, but they
58 only use the one overall grade at the top. Otherwise, it would become too complicated – they don’t think
59 there is a need to bother with all options for grading. She emphasizes that their choices on what to
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3 annotate are based on their own data. She also stresses that they used the medical literature to help them
4 decide what is important to look for in terms of annotating and embryo development. The embryologist
5 stresses that this particular model that they use could not be used in a different lab because it is based
6 on their data and also based on the media that they use. They've been using the same one for
7 approximately 2 years now. She says they are happy with the current model, but they could change it if
8 they wanted to. However, this cannot be done anytime, on the spot. It requires special permissions to
9 set up and should be done outside of the working day. Also, they are the ones who decide how much
10 weight to give certain embryo events. In the table of event scores they look at, I find out, the weight
11 assigned was determined by the lab staff. Therefore, even though TL gives them an embryo score, it is
12 determined by how they programmed the events to be weighed. (Author observation notes, Clinic C)
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17 While clinic C put significant effort into building a systematic TL process, not all clinics are
18 able to invest the same amount of resources into developing their algorithm. Research and
19 knowledge of statistical parameters is necessary to make optimal choices. All clinics
20 emphasized how their use of TL is particular to their situation and their expertise. Thus, how
21 TL is integrated largely depends on a seamless integration with the professional capabilities of
22 the lab where it is used. This discourse countered the wider enthusiasm for knowledge
23 standardization in embryo assessment through the use of TL. The study of local lab practices
24 (Latour 1983) reveals all the different ways in which TL can be used on the ground. We noticed
25 different levels of engagement with the algorithm functions of TL. Ultimately, staff always
26 have the option of using it simply as an incubator that generates embryo images. However, this
27 was not seen as cost-effective given the high cost of TL technologies. Ultimately, under the
28 current scientific lack of consensus around morphokinetic assessment, it is clinics that choose
29 if they want to engage in the process of algorithm creation/adaptation at all.
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49 **Choosing Embryos: Algorithmic Output and Questions of Expertise**

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51 Although TL algorithm scores are meant to improve embryo selection, we found that
52 this new knowledge dimension was not always easy to integrate within established professional
53 practices. Embryologists worked to incorporate this new technological expertise into their
54 routine, but also questioned the algorithmic output and how it might pose a challenge to
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3 professional expertise. During observations, we studied how TL algorithm scores are produced
4 and featured in clinical embryo transfer decisions. The types of scores produced by lab
5 algorithms varied slightly. For example, clinic D had a score that could go up to 75, while clinic
6 B had a score between 1 and 6. Others were receiving a letter grade output from the TL
7 algorithm. It is usually the highest score that indicates a good-quality embryo. Transferring
8 more than one embryo is discouraged in UK clinical practice with some exceptions, making
9 the task of choosing only one difficult, especially when there are several of good-quality.
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19 All clinics showed resistance to relying on TL scores exclusively when choosing
20 transfer embryos. Morphological grading is entrenched in decision-making. We observed
21 decisions being made on a case-by-case basis, according to professional judgement
22 (Greenhalgh et al. 2008). TL scores were sometimes viewed with skepticism or even ignored:
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28 The embryologist thinks that the score 65 embryo looks better than one with a score of 70, which are
29 meant to be 'better' according to the TL algorithm. She looks at the annotations and some annotation
30 scores appear in red when she looks at them in the table, meaning that the event did not happen within
31 the time expected. She wonders if maybe one of the staff did not annotate this properly. She suspects
32 that maybe a minor mistake was made because, to her, the 65 embryo looks better and she would choose
33 it over the higher scoring one. It is interesting that she attributes this to staff error rather than program
34 error. Also, the score doesn't seem to make her question her own judgement. (Author observation notes,
35 Clinic C)
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37 Here, the embryologist makes a judgement on embryo quality based on morphology.
38 For her, this overrides the TL score, which she suspects is lower because of erroneous
39 annotation. We witnessed a few such instances where morphology or the 'old scoring system'
40 was prioritized when choosing an implantation embryo. Often, staff felt more confident in the
41 established way of choosing embryos. This is not to say that embryologists do not care, more
42 generally, about TL data. It rather shows that standardizing the incorporation of new
43 information is rather difficult. Additionally, without clear evidence on the benefits of TL for
44 pregnancy rates, embryologists viewed the new scores with skepticism. Staff were aware of
45 the subjective dimensions of annotation and algorithm-creation. In the example above, a senior
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3 embryologist questioned the annotation process rather than her own judgement on the
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5 morphology of the embryo.
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8 Depending on their confidence regarding the TL score's robustness, professionals
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10 expressed being interested in using these scores. Their desire to do so, however, was limited to
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12 situations where many embryos of similar quality are available to choose from. Interestingly,
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14 re-watching the TL videos helped embryologists re-evaluate an embryo, if necessary. This
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16 enhanced confidence in transfer choices, independently of the algorithm feature. In our notes
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18 from clinic B, we wrote:
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21 I ask the embryologist what helps her decide is she's unsure which embryo to pick. She says she watches
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23 the videos side by side very slowly and looks for small anomalies (fragmentation %, for example she
24
25 says) and only after that she will look at the score that TL gives them. But she adds that the score should
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27 always be taken "with a grain of salt." She explains that they don't use it all that much (it is rarely
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29 necessary she seems to suggest). She's glad this patient's got many good embryos, but she says she
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31 won't need that many. (Author observation notes, Clinic B)

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33 Here the embryologist reinforces the need to be skeptical about TL scores. As we suggested
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35 above, this raises questions about the possibility of TL score standardization and the
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37 algorithm's authority in choosing embryos. However, as we already explained, algorithms can
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39 vary widely between clinics. Consequently, embryologists see TL integration as a local work-
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41 in-progress rather than knowledge passed through top-down standardized guidelines (Knaapen
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43 2014).

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45 We were also interested in how TL scores were deployed outside of the lab. TL scores
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47 were not usually discussed with patients, as they were seen as an element that might cause
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49 confusion. Furthermore, with no clear consensus on how to interpret them, staff felt it would
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51 be unwise to overemphasize these to patients. Scores were recorded by some clinics on patient
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53 forms, but more often than not, patients were only given the morphological embryo grade. As
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55 this classification system is more established, patients are able to search information about it
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57 online and decipher what it might mean in relation to implantation rates for that grade. As TL
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59 scoring can vary between labs, patients would find it harder to find relevant information on
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3 these scores. Nonetheless, embryologists explained to us that they do refer to TL videos in their
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5 conversations with patients as this allows them to explain what they have seen the embryo 'do.'

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8 TL facilitates the travel of embryo information outside of the lab. Thus, it creates the
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10 possibility, according to staff, of patients questioning their expertise and decisions on which
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12 embryos to transfer. Three clinics (B, C and D) offered patients the option to have a USB stick
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14 with the TL video of their implanted embryo. This option was not taken up very often, but
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16 some staff felt it could be better advertised. Regardless, the videos were usually shared only
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18 after a pregnancy was confirmed. Two clinics (A, E) in our sample avoided sharing TL images
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20 and videos, unless the patients brought this up themselves. Although not all, some
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22 embryologists felt anxious about the possibility of having their expertise questioned if patients
23
24 share TL videos with others who might provide a second opinion on their embryos.
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26 Consequently, sharing TL information, including potential access to an embryo livestream was
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28 often seen as an opportunity for the undermining of scientific expertise. This view resonated
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30 with many professionals:
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36 I think that's a difficult one because again it's their information but the problem will be is it's a very
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38 subjective field looking at embryos and you know this better than I do, I'm not a scientist, but it still
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40 subjective, there will still be some people that will still grade embryos slightly different to others
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42 although you have a pathway and follow protocol, there will be a slight variation and my worry is will
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44 people use it then and it has a negative effect. I want to take this to somebody else for a second opinion.
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46 And I think that's the only danger I see. Not that I don't think somebody should have a second opinion
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48 but it's a very, a subjective assessment and I know embryologists have pretty much now a
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50 standardization for grading of embryos but I still think that could happen. And taking that now to a
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52 private independent embryologists and you know, I just worry about the integrity of that. But I think
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54 it's powerful information, powerful but it's theirs, you know, it is their information but I just think it's
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56 powerful information that could be used sadly not always used in the right way. (Senior fertility nurse,
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58 Clinic B)
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61 Interestingly, our respondent emphasizes the limits of current standardization as it exists. She
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63 talks about the standardized morphological grading as it has been used for the past decades and
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65 contends that, even there, she sees issues around scientific objectivity. To a certain extent, the
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67 existence of TL and its functions threaten to disrupt the current order: the technology exposes
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69 patients to what was largely 'inside knowledge' before. As such, it becomes evident that

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3 underneath the surface of TL's algorithmic promises, lie uncertainties regarding the best use of
4 this technology and information generated through it. Enthusiasm for sharing TL videos with
5 patients varied significantly during our conversations with staff. Not only did each clinic have
6 different ways of providing patients with information, but each member of staff had differing
7 views on whether or not the process is beneficial for patients at all. The embryo information
8 that can be retrieved through TL was generally seen as having the potential to make patients
9 even more anxious about a process that is already challenging.
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21 **Conclusion**

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23 We have outlined above how local algorithmic practices co-exist in tension with
24 standardization expectations. Through the introduction of TL in IVF labs, professionals have
25 had to adapt to the demands of this new technology. The perceived subjective input that TL
26 requires deems the technology as an incomplete entity – an entity whose authority professionals
27 challenged periodically, while also working to improve algorithmic output. Through the
28 exploration of the TL case, we argue that biomedical algorithmic knowledge co-exists in
29 tension with complex lab routines and clinical contexts. This is partly a result of the input
30 needed from professionals to make the technology 'work' and the questions staff raise about
31 perceived subjective practices. STS scholars often conceptualize technologies as situated
32 (Aviles 2018; Coutard and Guy 2007). In the context of increased interest in the social life of
33 algorithms (Ziewitz 2015), we suggest that algorithms themselves can also be conceptualized
34 as situated practice. Moreover, we add evidence that actors may struggle with valuing
35 algorithmic technologies (Lee and Helgesson 2020) and that algorithms are not always
36 predictable (Neyland 2015). As seen in the TL case, the local embeddedness of algorithmic
37 practices impacts knowledge creation in ways that standardization efforts do not necessarily
38 anticipate prior to the introduction of the technology.
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3 Our findings show that embryologists working with TL are faced with numerous
4 decisions in relation to annotation processes, algorithm implementation, TL score use and how
5 to share TL information with patients. The analysis illuminates lab practices, thus
6 complementing previous studies of embryo selection (Helosvuori, 2019) and social science
7 studies of TL that focused on professionals' negotiation of EBM standards (Authors, XXXX),
8 visual characteristics of TL and its commercialization (Van de Wiel 2018, 2019). In local
9 practice, uncertainties around professional algorithmic input lead to TL disrupting and
10 rearranges professional practices, rather than straightforwardly resolving uncertainties in
11 embryo knowledge. We contend that consensus and standardization in embryo assessment are
12 ever-evolving processes and that TL has added increased complexities to this process rather
13 than having simplified it. Thus, we suggest that STS scholars pay attention to the disruptive
14 qualities of algorithmic technologies as they are used in biomedicine. We also suggest that the
15 degree of human machine interaction required by such technologies greatly shapes how they
16 are perceived by professionals. TL has raised questions about the authority of algorithmic
17 outputs and highlights how professional judgements feature the subjective/objective
18 dichotomy, where objectivity is associated with knowledge standardization and certainty, while
19 subjectivity is associated with a high level of human involvement.

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22 More broadly, our case study makes an important link between professional movements
23 encouraging increased knowledge standardization through algorithmic technologies and the
24 actual implementation of such standards (Greenhalgh et al. 2008; Knaapen 2014; Moreira,
25 2007; Timmermans and Berg 2003). As others have shown (Greenhalgh et al. 2008), clinical
26 decision-making is still a process that entails complexities that professionals have to navigate
27 based on local knowledge and their previous experience. Algorithmic standardization, in
28 particular, we suggest, is a process-in-the-making where the introduction of AI-based
29 technologies does not automatically lead to a straightforward generation of knowledge. As
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such, we stress the need to study algorithmic lab technologies at the local level, to understand: 1) how they reshape medical practice; 2) how the interplay between professional practice and such technologies shapes biomedical knowledge and 3) how algorithms and their output are incorporated and/or resisted in clinical practice.

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