Birth of Healthy Baby from 20-Year-Old Frozen Embryo Raises Ethical Questions

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Introduction

The Jones Institute for Reproductive Medicine at Eastern Virginia Medical School (Jones Institute) pioneered the first in-vitro fertilization (IVF) clinic in the United States in 1978. Three years later, in 1981, the Jones Institute reported the birth of the nation’s first IVF baby. Almost thirty years later, in 2010, the Jones Institute reported yet another milestone in reproductive medicine: the birth of a healthy baby boy from a 20-year-old frozen embryo.

The 20-year-old frozen embryo, along with others, had been created by an infertile couple who had undergone IVF treatment in 1990. The couple’s excess embryos were cryopreserved, or frozen at the pronuclear stage, and then placed in cryopreservation storage. After the couple was blessed with the birth of a baby boy through the IVF treatment, they subsequently decided in 1993 to anonymously donate their five remaining...
frozen embryos. The embryos were made available for donation, but remained in storage for an additional 16 years before they were finally matched with a 42-year-old woman, who had been undergoing infertility treatment for ten years.

Two of the five frozen 20-year-old embryos survived the thaw process and both were then transferred to the uterus of the 42-year-old recipient in August 2009. Only one embryo thrived in her womb, and in May 2010 after 41.5 weeks’ gestation, the 42-year-old woman delivered a healthy baby boy, weighing six pounds and 15 ounces.

Fertility specialists believe the 20-year-old embryo is the first of such age to produce a healthy baby. Until now, the previous record was held by two 13-year-old frozen embryos. In 2005, one 13-year-old frozen embryo produced a healthy baby girl in California, and in 2006, another 13-year-old embryo produced a healthy baby boy in Spain. Now, experts are claiming that frozen embryos may be stored for as long as 40 years. At this point, however, the oldest conceivable human frozen embryo available for IVF could be no more than 30-years-old since cryopreservation techniques were not used in human infertility treatment until the 1980s.

Millions of healthy children have been born through IVF, and the procedure is believed to be reasonably safe. Yet, fertility experts and researchers have long wondered whether IVF embryos—the embryos grown in laboratory petri dishes for several days—actually undergo subtle changes, and if so, whether those changes would cause adverse consequences to the health and development of the children born through IVF.

9 Dowling-Lacey, supra note 4, at 1.e1-1.e2.
10 Id.
11 Current Events, supra note 4.
12 As with the freezing process, the thawing process is carried out in multiple steps. See H.W. Michelmann and P. Nayudu, Cryopreservation of Human Embryos, 7 CELL AND TISSUE BANKING 135, 137-138 (2006), available at http://www.springlink.com/content/997700t16h72556g?fulltext.pdf.
13 Dowling-Lacey, supra note 4, at 1.e2.
14 Id. at 1.e2.
15 Id. at 1.e1.
18 Current Events, supra note 4.
19 Michelmann, supra note 12, at 135.
The news of the birth of the infant boy conceived from the 20-year-old embryo and the potential for 40-year-old embryos to emerge in the future now raise ethical questions that carry a host of medical, social, and psychological implications for the children born under such circumstances. Fertility specialists should be mindful of these ethical questions and implications and ensure that discussions concerning the same are incorporated in the informed consent conversations with their infertile patients who agree to use long-term frozen embryos.

**Medical Implications**

Embryos are frozen at a temperature of -196°C in liquid nitrogen with the use of a cryoprotectant agent. For many years, experts have speculated that frozen embryos stored at -196°C exist in a state of cryostability – meaning the embryos remain in suspended animation with no metabolic activity.

Since the inception of human embryo cryopreservation techniques in the 1980s, a paramount concern has been whether the cryopreservation procedure itself is safe. There is an ongoing concern that cryopreservation may pose medical risks to embryos and resulting children. Experts estimate that approximately 20 percent of embryos do not survive the freezing process, and the combination freezing/thawing processes potentially could cause embryo injury, genetic malformations or birth defects. But such adverse cryopreservation consequences must be clearly distinguished from the other potential risks associated with IVF that are currently being researched, such as certain birth defects, genetic disorders, or abnormal patterns in gene expression, all of which may be attributed to possible changes in the molecular biology of embryos grown in laboratory petri dishes.

During 2007 and 2008, researchers with the Jones Institute evaluated the impact of long-term cryostorage on post-thaw and implantation survival rates of frozen embryos. They analyzed computerized data from IVF patients who had used 11,768 frozen embryos.


23 *Cryopreservation and Storage*, supra note 6; see also Michelmann, supra note 12.
26 Michelmann, supra note 12, at 139.
27 Id.
28 See Kolata, supra note 21. Subtle changes in gene expression or in developmental patterns may not be obvious at birth and may not become apparent until adulthood, middle age, or old age.
29 Riggs, supra note 24, at 109.
between 1986 and 2007. The subject embryos had been frozen at the pronuclear and cleavage stages, with storage times ranging from 30 days to roughly 16 years. The researchers found that the post-thaw and implantation survival rates and the live birth rates for these pronuclear and cleavage stage embryos were similar regardless of the length of storage. After assuming all storage conditions were optimal for purposes of this study, the researchers concluded that the length of storage time had no significant effect on post-thaw survival rates or successful pregnancy outcomes.

Even though the Jones Institute researchers opine that frozen human embryos are relatively stable when stored at -196°C and that duration of storage has no obvious effect on pregnancy outcomes, these researchers, nevertheless, acknowledge the need for long-term follow-up of children born from frozen embryos. To date, there is no published data available on studies comparing the effects of long-term cryopreservation storage with normal childhood development. Normal childhood developmental milestones include gross motor, fine motor, cognitive, language, and social and emotional skills at age appropriate levels.

Of the limited studies that have been conducted on IVF-conceived children to date, the American Society for Reproductive Medicine (ASRM) and the Centers for Disease Control and Prevention (CDC) collectively reported the results of at least five studies evaluating behavior symptoms, academic achievement, birth weight, emotional health, and incidence of birth defects in IVF children. One study at the University of Pennsylvania compared the behaviors of IVF children with non-IVF children. In this study, the parents of IVF-children and the parents of non-IVF children were asked to complete The Child Behavior Checklist and a questionnaire about their children’s

30 Id. at 110.
31 The cleavage stage denotes embryos in the process of dividing or cleaving. This initially occurs on day 3 after fertilization. Riggs, supra note 24, at 110.
32 Id. at 110-111.
33 Id. at 113-114.
34 Id. at 112. From a practical standpoint, storage conditions may be adversely affected by repeatedly accessing storage tanks to retrieve embryos or by improperly maintaining storage tanks with liquid nitrogen.
35 Id. at 113-114.
37 ASRM is a multidisciplinary organization dedicated to the advancement of the art, science, and practice of reproductive medicine. Its mission is to pursue excellence in education and research and through advocacy on behalf of patients, physicians, and affiliated health care providers. See American Society of Reproductive Medicine, Mission Statement, available at http://www.asrm.org.
38 Highlights from ASRM09, supra note 22; Evaluation of Young Adults Conceived Via IVF, supra note 22; Grady, supra note 22 and Hitti, supra note 22.
39 The Child Behavior Checklist is a parent-report questionnaire on which the child is rated on various behavioral and emotional problems. It was developed by Thomas A. Achenbach and has been one of the widely used standardized measures in child psychology for evaluating maladaptive behavioral and
behavioral symptoms. Investigators analyzed the parents’ responses for 105 IVF children and 108 non-IVF children and concluded there were no significant differences in behavioral symptoms between the groups.

In the second study conducted at the University of Washington School of Medicine, researchers compared the birthweight and size of IVF newborns with spontaneously conceived non-IVF newborns and ultimately concluded there were no differences in their sizes at birth. These results were contrary to prior study results which suggested that IVF infants were smaller at birth and had more adverse outcomes than spontaneously conceived children. Now, it is believed that the underlying health of the parents may play a role in their offspring’s health regardless of the mode of conception.

In a third study, researchers at the University of Iowa used the Iowa Tests of Basic Skills to compare academic achievement test performance of IVF-conceived singletons, twins, and triplets between eight and 17 years of age. The researchers concluded that, although multiple gestations appear to have a slight detrimental effect on standardized test performance, the overall test scores of IVF children were high on the standardized scale.

In the fourth study, researchers evaluated and compared the first cohort of IVF-conceived young adults with their non-IVF counterparts and found the IVF young adults were happy and well-adjusted with no prevalence of increased susceptibility to chronic diseases. However, the researchers found there was a higher incidence of reports of clinical depression and attention deficit disorder/attention deficit hyperactivity disorder (ADD/ADHD) among the IVF young adults.

In the fifth study, researchers studied 281 women who underwent IVF treatment and found that their infants conceived through IVF were two to four times more likely to have certain birth defects than infants conceived naturally. The prevalent defect problems included heart problems, cleft lip and cleft palate, and esophageal and rectal

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40 Highlights from ASRM09, supra note 22.
41 Id.
42 Id.
43 Id.
44 Id.
45 Id.
46 These tests consist of standardized achievement batteries used in grades kindergarten through eighth grade. The tests cover several areas including mathematics, science, social studies, vocabulary, reading comprehension, word analysis, listening, and language skills. See University of Iowa College of Education, The Iowa Tests of Basic Skills, http://www.education.uiowa.edu.itp/itbs.
47 Id.
48 Id.
49 Evaluation of Young Adults Conceived Via IVF, supra note 22.
50 Id.
51 Grady, supra note 22 and Hitti, supra note 22.
abnormalities. However, it is not known whether IVF procedures increase the risk for these birth defects or whether infertility itself raises the risk for such defects.

None of these five studies mentioned above, however, distinguished between IVF children born from fresh embryos versus frozen embryos. It remains medically uncertain whether the processes of freezing and thawing embryos pose long-term medical and developmental risks to resulting children. Thus, there is a definitive need for long-term longitudinal research and studies comparing and/or contrasting the duration of human embryo cryopreservation storage with normal childhood development.

Social and Psychological Implications

Aside from the potential medical and developmental risks associated with embryo cryopreservation, ethics experts have also raised concerns about the potential social and psychological implications affecting the overall interests and well-being of children born from frozen embryos. Arguably, as the duration of embryo cryostorage periods lengthen, the adverse implications may exponentially increase, as well.

Consider for instance, the fact that the newborn baby boy born from the 20-year-old embryo and the son of the infertile couple who donated the 20 year-old embryo are biological siblings even though they were born 20 years apart. What if the non-biological parents of this newborn baby boy decide to tell him when he is older about the circumstances of his conception and the news that he has a brother?

The ASRM Ethics Committee supports the practice of parents disclosing to their children about the use of donor gametes in their conception, but ASRM also acknowledges the relative benefits of such disclosure are still under continued study. Disclosure is ultimately the parents’ choice. If, in this 20-year-old embryo case, the non-biological parents disclose to their son about his conception and genetic origins, how would this boy take such news? Would he be confused about his place in society and where he fits in his immediate family, given that he was born 20 years after conception? Would he experience identity problems and pursue the quest to find his biological roots only to learn that his parents were anonymous donors and he cannot have access to such information? Or worse yet, would he experience feelings of abandonment since he was left in storage for nearly 20 years before he was fortuitously matched with his current

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52 Id.
53 Id.
54 Michelmann, supra note 12.
57 Id. at 527. When parents disclose such information, there has been a push in recent years to allow children access to non-identifying information about the gamete donors if they request it and access to identifying information about donors if all parties agree. Id.
58 Potentially, a child born from a frozen embryo may experience identity problems regardless of the length of cryopreservation storage. See Winslade, supra note 55.
parents? The psychological interests and well-being of the children (and later adults) born from long-term frozen embryos cannot and should not be ignored.

Conclusion

IVF patients who choose to use long-term frozen embryos, whether their own or those donated from others, must consider the medical risks and social and psychological implications for their children produced from these embryos. Fertility specialists should be especially cognizant of these risks and implications and should thoroughly discuss them accordingly with their patients during the informed consent processes. Moreover, fertility specialists should push for long-term longitudinal research and studies evaluating the impact of long-term frozen embryo storage on normal childhood developmental skills and adult emotional and psychological development.
A woman in Tennessee has given birth to a baby daughter that was frozen as an embryo 24 years ago. It is believed to be a world record for frozen embryo to result in a successful birth. Emma was implanted into her mother's uterus earlier this year through a process called frozen embryo transfer. After a successful pregnancy, baby Emma was born with a minimum of fuss and her parents couldn't be happier. "Emma is such a sweet miracle," Dad Benjamin said. "I think she looks pretty perfect to have been frozen all those years ago." A woman has just recently given birth to a child born from the longest known frozen human embryo. Tina Gibson, from Eastern Tennessee, gave birth to Emma Wren Gibson on November 25th. The event may break a new record, while simultaneously raising questions about what to do with preserved embryos. Gibson's successful pregnancy and delivery likely beat out the previous record for a successful birth from a frozen embryo, which was an embryo frozen for 20 years. The embryo that became Emma was initially frozen back in October 1992, when Tina was only 18 months old. Tina noted that had Emma been born at the initial time, they could have easily been the right age to be friends. ADVERTISEMENT.