### Chapter 10

# The Art of Ethical Dimensions in Stem Cell Research

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## A. Significance of Stem Cell Research

Stem cell research is a groundbreaking field with vast potential to revolutionize medicine and biotechnology through regenerative therapies and disease treatments (Mao et al., 2022). Stem cell research, promising for medicine and regeneration, faces ethical dilemmas, mainly concerning embryonic cells. Stem cell advancements, like CRISPR-Cas9 gene editing, offer transformative prospects for personalized medicine (Wang et al., 2022). Stem cells hold immense transformative potential in regenerative medicine, disease modeling, and drug development, offering hope for treating various debilitating conditions.

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Responsible research practices in stem cell studies require collaboration among policymakers, researchers, bioethicists, and the public to balance progress with ethics, ensuring benefits and safeguarding stakeholder well-being (Neil et al., 2022). Ethical issues in stem cell researches, from the moral status of embryos to stem cell sourcing, demand careful attention. Upholding informed consent, ethical reviews, and standards are essential to safeguard the rights and well-being of both human and animal participants. (Lovell-Badge, et al., 2021). As stem cell therapies advance to clinical trials and commercialization, strong legal frameworks are essential to guarantee patient safety, oversee research, and avoid exploitation (Moradi et al., 2019). Ultimately, the vast benefits of stem cell research must be balanced with its ethical considerations to align science with societal values.

### B. Ethical Issues in Stem Cell Research

Ethics in scientific research is a fundamental aspect that governs the conduct of scientists and researchers in their pursuit of knowledge and advancements (Sato & Suzuki, 2022). At its core, ethics in scientific research involves adherence to moral principles, values, and standards that ensure the responsible and respectful treatment of all individuals involved, including research participants, colleagues, and the broader community. Ethical considerations in stem cell research are particularly critical due to the complex nature of these cells and their potential implications for human health (Assen et al., 2021).

Central to the concept of ethics in stem cell research is the principle of respect for human dignity and autonomy. This principle necessitates obtaining informed consent from research participants, ensuring they are fully aware of the risks, benefits, and potential outcomes of the study. Researchers must also protect the privacy and confidentiality of participants, respecting their rights to autonomy and self-determination throughout the research process (Harris et al., 2022).

Another crucial ethical consideration in stem cell research relates to the source of stem cells. The use of embryonic stem cells (ESCs) has been a subject of significant debate due to the destruction of embryos in the process of obtaining these cells. This raises complex questions regarding the moral status of embryos and the balance between scientific advancement and respect for human life. As a result, researchers have sought alternative sources of stem cells, such as induced pluripotent stem cells (iPSCs), which can be generated from adult somatic cells without the ethical concerns associated with embryonic sources (Singh et al., 2015).

Moreover, researchers must uphold principles of scientific integrity and honesty, ensuring transparent reporting of research findings. Negative results and potential limitations of the research should be openly communicated, preventing publication bias and fostering a more comprehensive understanding of stem cell science (Valdés & Lecaros, 2023).

Ethical issues in stem cell research also extend to the potential for commercialization and patenting of stem cell technologies. The pursuit of financial gain can sometimes conflict with considerations of equitable access to therapies and the public good. Striking a balance between incentivizing innovation and ensuring affordability and accessibility of stem cell treatments is essential to navigate these ethical challenges (Alahmad et al., 2020).

Stem cells hold immense promise for regenerative medicine, but the ethical considerations surrounding their sources have been a focal point of debate in the scientific community and beyond (Volarevic et al., 2018). Among the various types of stem cells, embryonic stem cells (ESCs) derived from early-stage embryos have been at the heart of controversy due to ethical implications related to the moral status of embryos and the potential for harm. Understanding the arguments for and against the use of embryonic stem cells is crucial for a comprehensive analysis of the ethical challenges in stem cell research (Al-Agele, 2023).

### Harnessing Embryonic Stem Cells: Unveiling their Potential

Embryonic stem cells offer a fascinating discussion due to their unique features, especially pluripotency. This trait allows these cells to transform into various cell types, forming the complex human body. Such potential brings optimism to regenerative medicine and disease research, paving the way for new medical breakthroughs (Żakowska-Henzler et al., 2023).

Pluripotency in embryonic stem cells holds transformative promise for regenerative therapies. Supporters see a future where regrowing damaged tissues, like heart muscle after an attack or nerve cells in degenerative diseases, is possible, challenging traditional medical boundaries (Cho et al., 2022).

Moreover, embryonic stem cells are valued not only for their adaptability but also for their ability to self-renew indefinitely. This trait provides a consistent supply for research, meeting the growing demand in scientific studies and ensuring continuous exploration (Chen et al., 2022; Varzideh et al., 2023).

Embryonic stem cell advocates navigate beyond science to ethics, addressing questions about life's beginnings and research morality. Nonetheless, they believe the research's potential could transform healthcare and offer healing to many (Hauskeller et al., 2019).

Embryonic stem cells represent a complex potential in science, with their pluripotency offering vast possibilities and their self-renewal fueling continuous knowledge. Their significance triggers both scientific and ethical discussions. Navigating this, a balanced approach to ethics, research, and hope is essential for progress (Blasimme & Sugarman, 2023).

### 2. Arguments for the Use of Embryonic Stem Cells

Advocates for embryonic stem cells highlight their pluripotency, which lets them transform into any human cell type (Fléchon, 2022). Embryonic stem cells' ability to differentiate offers potential

for regenerative therapies and disease modeling. Their prolonged self-renewal also makes them a sustainable research resource (Chen et al., 2022).

### 3. Arguments Against the Use of Embryonic Stem Cells

A main ethical concern with embryonic stem cells is the embryo's destruction during extraction. Critics believe embryos, with their potential for human development, warrant protection, equating the process to taking human life and raising ethical issues about their use in research (Koplin, 2023).

The ethical implications of obtaining and using embryonic stem cells extend beyond their origin. Their potential clinical applications introduce additional concerns, such as the risk of teratoma formation—the development of tumors comprising various cell types—following transplantation. Researchers must carefully weigh the potential benefits of embryonic stem cell-based therapies against these risks and address safety concerns responsibly (Volarevic et al., 2018).

As stem cell research advances, the ethical discussions about their source and use highlight the need for a careful approach (Abubakar et al., 2023). Balancing science with ethics and exploring alternatives like iPSCs and adult stem cells is crucial for advancing stem cell research while maintaining societal values and standards (Moradi et al., 2019).

# C. Informed Consent and Human Subjects in Stem Cell Studies

Informed consent is fundamental to ethical research, ensuring participants fully understand a study's aims, risks, benefits, and their rights. In stem cell research, with its potential experimental treatments, securing informed consent is vital to protect participants' autonomy and well-being (Riordan et al., 2022).

Stem cell studies, from lab research to clinical trials with humans, require researchers to offer clear, comprehensible information

to potential participants or their legal representatives (Yui et al., 2022). Informed consent should promote open dialogue, ensuring participants' voluntary, and informed choices. Consent should be continuous, with the option to withdraw at any time without consequences (Padilla et al., 2022).

Particular attention is required when obtaining informed consent for stem cell clinical trials, as these studies involve novel therapies with unknown risks and potential benefits. Participants must be thoroughly informed about the investigational nature of the treatment and the uncertainties surrounding its outcomes (van Rijssel et al., 2022). Researchers must uphold the principles of beneficence and non-maleficence, ensuring that the potential benefits outweigh the risks for research participants (Hosseini et al., 2022).

In the context of stem cell research, the role of human subjects and human donors is pivotal. Both aspects require rigorous ethical considerations, particularly concerning informed consent, the fate of the cells collected, storage and potential use of these cells, and compensation to donors, including considerations around commercialization and potential royalties (Assen et al., 2021).

### Human Subjects in Stem Cell Research

Human subjects participating in stem cell research, especially in clinical trials, need to be fully informed about the investigational nature of the study. This includes a clear understanding of the potential risks and benefits, the experimental status of the treatments, and the fact that outcomes are uncertain. Continuous informed consent is crucial, and participants must have the option to withdraw at any time without any repercussions. This aligns with ethical principles, ensuring the autonomy and safety of participants (Yamanaka, 2020; Zarzeczny, 2019).

#### 2. Human Donors in Stem Cell Research

In cases where stem cells are sourced from human donors, informed consent becomes even more complex. Donors must understand the purpose of the donation, the process involved in cell collection, and the future use of these cells (Guo et al., 2021).

#### Fate of the Donated Cells

Based on the purposes, the fate of the donated cells will fall into two categories.

- Research purposes: Initially, cells might be collected for specific research purposes. This could involve studying cell behavior, testing new treatments, or understanding disease mechanisms (Charitos et al., 2021).
- 2) Post-research fate: After the research is completed, the cells might be either destroyed, stored for future research, or used for other research projects. The specific fate of the cells should be transparently communicated to the donor during the consent process (Orzechowski et al., 2021).

### b. Storage and Use of Donated Cells

Donated cells may be stored in biobanks. The duration of storage can vary depending on the research requirements and the viability of the cells. These cells could be used for (Zalaf et al., 2020):

- 1) future research initiatives;
- development of treatments for other individuals, subject to ethical approvals and regulatory guidelines;
- 3) educational or training purposes in some cases.

### c. Compensation to Donors

The issue of compensation for donors is complex. Typically, donors are not paid for their donations as it could influence their decision to donate and raise ethical concerns about commodification of body parts. However, donors may be compensated for associated costs like travel or time off work (Zalaf et al., 2020).

### 3. Commercialization and Royalties

If stem cells are commercialized, the ethical and legal frameworks become even more intricate. Generally, donors do not receive royalties from products developed from their cells. The principle here is that once donated, the cells become a part of a common resource for scientific advancement. However, this is a topic of ongoing ethical debate, particularly in cases where the commercial use of donated cells leads to significant profit (Padovano et al., 2022).

Ultimately, both human subjects and donors in stem cell research are protected by stringent ethical guidelines, emphasizing informed consent, the careful handling and use of donated cells, and clear communication about all aspects of the research and its potential implications. The principles of autonomy, beneficence, non-maleficence, and justice, guide these practices, ensuring ethical conduct in this rapidly evolving field (Lovell-Badge, et al., 2021).

# D. Considerations for Conducting Research on Adult and Somatic Stem Cells

Amidst the ethical debates surrounding embryonic stem cells, adult and somatic stem cells offer a promising alternative for research and therapeutic applications. Adult stem cells are found in various tissues throughout the body and play a crucial role in tissue repair and regeneration. Their inherent ability to self-renew and differentiate into specialized cell types has sparked interest in exploring their therapeutic potential. However, conducting research on adult and somatic stem cells raises unique ethical considerations that must be thoughtfully addressed (Hoang et al., 2022).

One ethical consideration involves the collection of adult stem cells, which can be obtained from donors through minimally invasive procedures. Ensuring that the collection process poses minimal risk to the donor while adhering to the principles of informed consent and privacy is essential (Escoto et al., 2023). Researchers must also consider issues of equity and justice, as access to stem cell therapies

should not be limited to a privileged few but should be accessible to a diverse range of patients, regardless of socioeconomic status or geographical location (Farajkhoda, 2017).

In addition to the sourcing of adult stem cells, ethical questions arise concerning the use of somatic stem cells for research purposes. Somatic stem cells are responsible for maintaining and renewing specific tissues in the body, and their exploration offers insights into tissue-specific diseases and regenerative medicine (Ji et al., 2023). Ethical considerations revolve around potential risks associated with manipulating and modifying somatic stem cells, as well as ensuring their safe application in clinical trials and therapies (de Jongh et al., 2022).

# E. Ethical Dilemmas in Stem Cell Clinical Trials and Experimental Therapies

The translation of stem cell research into clinical trials and experimental therapies presents ethical dilemmas. Rigorous ethical standards must be upheld to protect research participants and ensure the validity of findings. Ethical concerns involve patient selection, informed consent, and transparent reporting of outcomes (Drolet et al., 2023). Offering unregulated stem cell treatments outside clinical trials raises safety and exploitation concerns, necessitating comprehensive guidelines and regulations. Responsible and ethical practices are crucial to fulfill the promise of stem cell research while upholding human rights and scientific integrity (Peng et al., 2020).

### F. Ethical Guidelines and Best Practices

Ethical guidelines serve as critical foundations for ensuring the responsible conduct of stem cell research. Leading organizations, including the International Society for Stem Cell Research (ISSCR), the National Institutes of Health (NIH), and the World Health Organization (WHO), have developed comprehensive frameworks to address the ethical complexities of stem cell research and its

applications. These guidelines cover essential aspects such as informed consent, protection of research participants, transparency in scientific reporting, and the appropriate use of stem cell sources. By providing a shared ethical framework, these guidelines facilitate collaboration among researchers, promote transparency, and instill public confidence in stem cell science (Assen et al., 2021).

As stem cell research continues to advance, it is essential to develop a robust framework for ethically responsible practices. This framework must embrace ethical considerations at every stage of the research process, from laboratory investigations to clinical applications. Researchers and institutions should prioritize the principles of beneficence, non-maleficence, autonomy, and justice in their work, ensuring that the potential benefits of stem cell research are maximized while minimizing any potential harm or risk to patients and society (Kidha, 2020).

Central to this framework is the establishment of transparent and accountable mechanisms for oversight. Ethical review boards and institutional committees play a vital role in evaluating proposed research studies, scrutinizing ethical aspects, and ensuring that research adheres to the established guidelines and regulations. Engaging with stakeholders, including patients, the public, and ethicists, can further enrich the ethical considerations of stem cell research, facilitating diverse perspectives and avoiding undue concentration of decision-making power (Assen et al., 2022).

The pursuit of scientific progress and ethical considerations in stem cell research is an intricate balancing act. On one hand, the urgency to address critical health challenges propels researchers to explore novel avenues and potential therapeutic applications. On the other hand, responsible scientific inquiry must be guided by robust evidence, adherence to ethical guidelines, and patient safety (Weinbaum et al., 2019).

Researchers should conduct stem cell research with humility, dedication to thorough study design, and peer review. Transparently sharing all results, good or bad, supports scientific integrity. Reviewing

the ethics of new technologies, like gene editing, ensures alignment with societal values and priorities (Joseph et al., 2022).

Ethical training and education are indispensable for researchers and professionals engaged in stem cell research. As new technologies and discoveries unfold, staying abreast of ethical guidelines and best practices is essential for informed decision-making. Researchers should receive comprehensive training that addresses the nuances of informed consent, patient privacy, the responsible use of stem cell sources, and the potential implications of their work on patients, communities, and future generations (Assen et al., 2022).

Promoting ethical reflection and open discussions in research institutions is crucial. Interdisciplinary collaborations and dialogues among researchers, ethicists, and policymakers ensure ethics remain central to stem cell research (Torres-Padilla et al., 2020).

# G. Ethical Guidelines and Best Practices in Indonesia

Indonesia, a diverse archipelago, blends cultural spirituality, religious jurisprudence, and Western bioethics. Medical professionals in Indonesia must balance international practices with these unique cultural perspectives (Hefner, 2021). In Indonesian bioethics, personal autonomy and informed consent are paramount. While aligning with international standards, Indonesia emphasizes respecting cultural and spiritual beliefs in the decision-making process for medical or research activities (Subandi et al., 2023). Indonesian ethics prioritize collective well-being. While individual rights are essential, they might be limited during public health crises. Any such measures should be proportionate, transparent, and temporary to prevent potential authoritarianism (Casey & Vermeule, 2022). In Indonesia, biomedical research follows stringent ethical rules. Human cloning is banned, as are genetic modifications inheritable by future generations, reflecting global caution and Indonesia's respect for human life (Florea, 2023; Staunton et al., 2019). In Indonesia, end-of-life care is influenced by diverse religious beliefs. Physicians must understand these cultural nuances, especially regarding topics like passive euthanasia, ensuring alignment with patients' spiritual values and family wishes (Kuhn, 2022). In Indonesia, the challenge for policymakers is translating ethical norms into laws. Due to the evolving nature of medicine and ethics, regular reviews involving bioethicists, religious leaders, and the public are essential (Ten Have & Neves, 2021). In Indonesia, ethical guidelines merge international standards with local cultures and beliefs. For healthcare and research success, it's vital to blend global insights with an understanding of the unique Indonesian context (Charitos et al., 2021; Fauziah & Mukhlis, 2019).

In the Indonesian context, the practical application of ethical guidelines in healthcare and biomedical research often reflects a unique blend of international standards and local cultural and religious values. This duality presents both challenges and opportunities in implementing ethical practices (Mathur et al., 2019).

#### 1. Practical Realities in Indonesia

### Balancing Autonomy and Collective Well-being

Indonesian healthcare professionals often navigate a fine line between respecting individual autonomy and prioritizing collective well-being. This balance becomes particularly evident during public health emergencies, where individual rights might be tempered for the greater good. The response to the COVID-19 pandemic is a case in point, where the government had to implement measures that balanced individual freedoms with public health necessities (Newnham & Kirkham, 2019; Søvold et al., 2021).

### b. Cultural Sensitivity in Medical Decision-Making:

In a culturally diverse nation like Indonesia, medical decisions are heavily influenced by the patient's cultural and religious background. This sensitivity is especially apparent in end-of-life care, where healthcare providers must align their practices with the patient's religious beliefs and family preferences, which can vary significantly across the archipelago (Pentaris & Christodoulou, 2021).

#### c. Bioethics in Research

In biomedical research, Indonesian researchers adhere to stringent ethical guidelines that prohibit practices like human cloning and inheritable genetic modifications. These regulations reflect a global cautious approach and are in line with Indonesia's respect for the sanctity of human life. The challenge lies in ensuring these guidelines are consistently applied and understood across various research institutions, which can be diverse in their resources and expertise (Earp et al., 2020).

### d. Regulatory Framework

Indonesia's approach to bioethics in medicine and research is to harmonize international norms with local beliefs and practices. However, translating these ethical norms into enforceable laws is an ongoing challenge. The dynamic nature of medical science requires that these laws and guidelines be regularly reviewed and updated, involving a wide range of stakeholders including bioethicists, religious leaders, and the public (Pimenta et al., 2021).

### e. Education and Training

To effectively implement these ethical guidelines, there is a need for continuous education and training for medical professionals and researchers. This training should not only cover the technical aspects of bioethics but also emphasize the importance of cultural competence in dealing with patients from diverse backgrounds (Ignatowicz et al., 2022).

### d. Public Awareness and Engagement

Public understanding and engagement in bioethical issues are crucial for the success of healthcare policies in Indonesia. This involves educating the public about their rights and the ethical considerations in medical treatments and research, thus fostering a more informed and participative society (Fletcher, 2023).

Ultimately, the reality of applying ethical guidelines in Indonesia's healthcare and research sectors involves a complex interplay of respecting international standards while also honoring the nation's diverse cultural and religious ethos. Continuous education, regulatory vigilance, and public engagement are key to navigating these challenges effectively (Resosudarmo, 2022).

### H. Future Directions and Ethical Deliberations

As stem cell research advances, new technologies continue to emerge, each with their unique ethical implications. One such technology is organoid development, which involves growing three-dimensional mini organs in vitro, providing a platform for studying human organ development and disease. While organoids hold great promise for disease modeling and drug testing, concerns arise over the potential for these structures to acquire unforeseen functions or even consciousness, raising questions about the ethical status of organoids and the boundaries of human-like entities (Kendal, 2022).

Additionally, while these technologies offer potential benefits in personalized medicine and regenerative therapies, ethical considerations must address issues of consent, equity, and the responsible use of powerful gene editing tools (Li et al., 2023). As these emerging technologies push the boundaries of what is scientifically possible, the ethical dimensions of responsible innovation must be carefully deliberated (Stahl et al., 2021).

The development of regenerative medicine and tissue engineering holds tremendous potential for addressing critical health challenges, such as organ failure and tissue degeneration. By harnessing the regenerative capabilities of stem cells, researchers aim to create replacement tissues and organs tailored to individual patients, minimizing the risk of rejection and enabling more effective treatments (Dzobo et al., 2018).

However, ethical considerations accompany these advancements. As researchers seek to create complex human tissues or organs using stem cells and scaffold materials, questions arise about the moral status

of these engineered tissues. Determining when these structures attain the status of a living organism or individual requires thoughtful ethical deliberation (Chen & Liu, 2016). The potential for commodification and commercialization of human body parts further complicates the ethical landscape, calling for robust safeguards to uphold the dignity of the human body and protect patients from undue exploitation (de Kanter et al., 2023; Lau, 2023).

### I. Conclusion

Stem cell research offers vast medical potential, but poses complex ethical challenges, from embryonic stem cell use to informed consent and equitable access. The legal landscape features intricate regulations and patents affecting research and affordability. Ensuring a responsible approach requires collaboration among researchers, ethicists, patients, and the public, supported by ethical boards and global partnerships. As the field advances, continuous dialogue, ethical awareness, and responsible innovation are vital to align scientific progress with societal values and well-being.

### References

- Abubakar, M., Masood, M. F., Javed, I., Adil, H., Faraz, M. A., Bhat, R. R., Fatima, M., Abdelkhalek, A. M., Buccilli, B., & Raza, S. (2023). Unlocking the mysteries, bridging the gap, and unveiling the multifaceted potential of stem cell therapy for cardiac tissue regeneration: A narrative review of current literature, ethical challenges, and future perspectives. *Cureus*, *15*(7), Article e41533. https://doi.org/10.7759/cureus.41533
- Al-Agele, R. A. A. (2023). Considering the safety of embryonic stem cells for medical use while ignoring any ethical concerns: A review. *Diyala Journal for Veterinary Science*, 1(1), 126–142. https://djvs.uodiyala.edu.iq/index.php/djvs/article/view/89
- Alahmad, G., Aljohani, S., & Najjar, M. F. (2020). Ethical challenges regarding the use of stem cells: Interviews with researchers from Saudi Arabia. BMC Medical Ethics, 21(1), Article 35. https://doi.org/10.1186/ s12910-020-00482-6

- Assen, L. S., Jongsma, K. R., Isasi, R., Tryfonidou, M. A., & Bredenoord, A. L. (2021). Recognizing the ethical implications of stem cell research: A call for broadening the scope. *Stem Cell Reports*, *16*(7), 1656–1661. https://doi.org/10.1016/j.stemcr.2021.05.021
- Assen, L. S., Jongsma, K. R., Isasi, R., Tryfonidou, M. A., & Bredenoord, A. L. (2022). Roles and responsibilities in stem cell research: A focus group study with stem cell researchers and patients. *Regenerative Medicine*, *17*(7), 445–459. https://doi.org/10.2217/rme-2022-0019
- Blasimme, A., & Sugarman, J. (2023). Human stem cell-derived embryo models: Toward ethically appropriate regulations and policies. *Cell Stem Cell*, 30(8), 1008–1012. https://doi.org/10.1016/j.stem.2023.06.007
- Casey, C., & Vermeule, A. (2022). Myths of common good constitutionalism [Working paper no. 22-09]. *Harvard Journal of Public Policy*, 45(1), 103–146. https://ssrn.com/abstract=4030763
- Charitos, I. A., Ballini, A., Cantore, S., Boccellino, M., Di Domenico, M., Borsani, E., Nocini, R., Di Cosola, M., Santacroce, L., & Bottalico, L. (2021). Stem cells: A historical review about biological, religious, and ethical issues. Stem Cells International, 2021, Article 9978837. https:// doi.org/10.1155/2021/9978837
- Chen, F. M., & Liu, X. (2016). Advancing biomaterials of human origin for tissue engineering. *Progress in Polymer Science*, *53*, 86–168. https://doi.org/10.1016/j.progpolymsci.2015.02.004
- Chen, G., Yin, S., Zeng, H., Li, H., & Wan, X. (2022). Regulation of embryonic stem cell self-renewal. *Life (Basel)*, *12*(8), Article 1151. https://doi.org/10.3390/life12081151
- Cho, S., Discher, D. E., Leong, K. W., Vunjak-Novakovic, G., & Wu, J. C. (2022). Challenges and opportunities for the next generation of cardiovascular tissue engineering. *Nature Methods*, 19(9), 1064–1071. https://doi.org/10.1038/s41592-022-01591-3
- de Jongh, D., Massey, E. K., & Bunnik, E. M. (2022). Organoids: A systematic review of ethical issues. *Stem cell research & therapy*, 13(1), Article 337. https://doi.org/10.1186/s13287-022-02950-9
- de Kanter, A. F. J., Jongsma, K. R., Verhaar, M. C., & Bredenoord, A. L. (2023). the ethical implications of tissue engineering for regenerative purposes: A systematic review. *Tissue Engineering Part B: Reviews*, 29(2), 167–187. https://doi.org/10.1089/ten.TEB.2022.0033
- Drolet, M.-J., Rose-Derouin, E., Leblanc, J.-C., Ruest, M., & Williams-Jones, B. (2023). Ethical Issues in research: Perceptions of researchers,

- research ethics board members and research ethics experts. *Journal of Academic Ethics*, 21(2), 269–292. https://doi.org/10.1007/s10805-022-09455-3
- Dzobo, K., Thomford, N. E., Senthebane, D. A., Shipanga, H., Rowe, A., Dandara, C., Pillay, M., & Motaung, K. S. C. M. (2018). Advances in regenerative medicine and tissue engineering: Innovation and transformation of medicine. *Stem Cells International*, 2018, Article 2495848. https://doi.org/10.1155/2018/2495848
- Earp, B. D., Demaree-Cotton, J., Dunn, M., Dranseika, V., Everett, J. A. C., Feltz, A., Geller, G., Hannikainen, I. R., Jansen, L. A., Knobe, J., Kolak, J., Latham, S. R., Lerner, A. S., May, J., Mercurio, M. R., Mihailov, E., Rodríguez-Arias, D., Rodríguez López, B., Savulescu, J., ... & Tobia, K. P. (2020). Experimental philosophical bioethics. AJOB Empirical Bioethics, 11(1), 30–33. https://doi.org/10.1080/23294515.2020.1714792
- Escoto, M., Issa, F., Cayouette, F., Consolo, H., Chaudhury, P., Dhanani, S., Jiang, W., Oniscu, G. C., Murphy, N., Rockell, K., Weiss, M. J., & Dieudé M. (2023). Research and innovation in organ donation: Recommendations from an international consensus forum. *Transplantation Direct*, *9*(5), Article e1446. https://doi.org/10.1097/txd.0000000000001446
- Farajkhoda, T. (2017). An overview on ethical considerations in stem cell research in Iran and ethical recommendations: A review. *International Journal of Reproductive Biomedicine*, 15(2), 67–74.
- Fauziah, R. F., & Mukhlis, M. F. (2019). Embryonic stem cells in stroke treatment health laws and shariah perspective. *International Journal of Innovation, Creativity and Change*, 6(1), 362–376.
- Fléchon, J.-E. (2022). What are ES cells? In L. M. Houdebine (Ed.), *Transgenic Animals* (157–166). CRC Press.
- Fletcher, F. E. (2023). Ethical, legal, and social implications of genomics research: Implications for building a more racially diverse bioethics workforce. *The American Journal of Bioethics*, 23(7), 106–108. https://doi.org/10.1080/15265161.2023.2207519
- Florea, M. (2023). Withdrawal of consent for processing personal data in biomedical research. *International Data Privacy Law*, *13*(2), 107–123. https://doi.org/10.1093/idpl/ipad008
- Guo, R., Mu, W., Liu, X., Zhang, J., Liu, B., Du, X., He, J., Ma, J., & Cui, H. (2021). Reprogramming of a human induced pluripotent stem cell

- line from one 48-year-old healthy male donor. *Stem Cell Research*, 53, Article 102339. https://doi.org/10.1016/j.scr.2021.102339
- Harris, A. R., Walker, M. J., & Gilbert, F. (2022). Ethical and regulatory issues of stem cell-derived 3-dimensional organoid and tissue therapy for personalised regenerative medicine. *BMC medicine*, *20*(1), Article 499. https://doi.org/10.1186/s12916-022-02710-9
- Hauskeller, C., Manzeschke, A., & Pichl, A (Eds.). (2019). The matrix of stem cell research: An approach to rethinking science in society. Routledge.
- Hefner, R. W. (2021). Islam and institutional religious freedom in Indonesia. *Religions*, *12*(6), Article 415. https://www.mdpi.com/2077-1444/12/6/415
- Hoang, D. M., Pham, P. T., Bach, T. Q., Ngo, A. T., Nguyen, Q. T., Phan, T. T., Nguyen, G. H., Le, P. T., Hoang, V. T., Forsyth, N. R., Heke, M., & Nguyen, L. T. (2022). Stem cell-based therapy for human diseases. Signal transduction and targeted therapy, 7(1), Article 272. https://doi.org/10.1038/s41392-022-01134-4
- Hosseini, A., Jackson, A., & Bahramnezhad, F. (2022). Ethical considerations in interventional studies: A systematic review. *Acta Medica Iranica*, 60(10), 609–614. https://doi.org/10.18502/acta.v60i10.11550
- Ignatowicz, A., Slowther, A. M., Bassford, C., Griffiths, F., Johnson, S., & Rees, K. (2022). Evaluating interventions to improve ethical decision making in clinical practice: a review of the literature and reflections on the challenges posed. *Journal of Medical Ethics*, 49(2), 136–142. https://doi.org/10.1136/medethics-2021-107966
- Ji, S., Xiong, M., Chen, H., Liu, Y., Zhou, L., Hong, Y., Wang, M., Wang, C., Fu, X., & Sun, X. (2023). Cellular rejuvenation: Molecular mechanisms and potential therapeutic interventions for diseases. *Signal transduction* and targeted therapy, 8(1), Article 116. https://doi.org/10.1038/s41392-023-01343-5
- Joseph, A. M., Karas, M., Ramadan, Y., Joubran, E., & Jacobs, R. J. (2022). Ethical perspectives of therapeutic human genome editing from multiple and diverse viewpoints: A scoping review. *Cureus*, *14*(11), Article e31927. https://doi.org/10.7759/cureus.31927
- Kendal, E. (2022). Ethical, legal and social implications of emerging technology (ELSIET) symposium. Journal of Bioethical Inquiry, 19(3), 363–370. https://doi.org/10.1007/s11673-022-10197-5

- Kidha, D. K. (2020). *Human embryonic stem cell research in transplantation and regenerative medicine: A principlist assessment* [Thesis]. Stellenbosch University. http://hdl.handle.net/10019.1/107871
- Koplin, J. J. (2023). Response to the ISSCR guidelines on human–animal chimera research. *Bioethics*, *37*(2), 192–198. https://doi.org/10.1111/bioe.13104
- Kuhn, E. (2022). Why me?-The concept of physicians' spiritual self-care: A contribution to professional and organisational ethics. In A. H. Seidlein & S. Salloch (Eds.), *Ethical challenges for healthcare practices at the end of life: Interdisciplinary perspectives* (55–72). Springer. https://doi.org/10.1007/978-3-030-83186-8\_4
- Lau, P. L. (2023). Evolved eugenics and reinforcement of "Othering": renewed ethico-legal perspectives of genome editing in reproduction. *BioTech*, *12*(3), Article 51. https://doi.org/10.3390/biotech12030051
- Li, S., Tang, H., Li, C., Ma, J., Ali, M., Dong, Q., Wu, J., Hui, Y., & Sun, C. (2023). Synthetic biology technologies and genetically engineering strategies for enhanced cell therapeutics. *Stem Cell Reviews and Reports*, 19(2), 309–321. https://doi.org/10.1007/s12015-022-10454-5
- Lovell-Badge, R., Anthony, E., Barker, R. A., Bubela, T., Brivanlou, A. H., Carpenter, M., Charo, R. A., Clark, A., Clayton, E., & Cong, Y. (2021). ISSCR guidelines for stem cell research and clinical translation: The 2021 update. *Stem Cell Reports*, 16(6), 1398–1408. https://doi.org/10.1016/j.stemcr.2021.05.012
- Mao, J., Saiding, Q., Qian, S., Liu, Z., Zhao, B., Zhao, Q., Lu, B., Mao, X., Zhang, L., Zhang, Y., Sun, X., & Cui, W. (2022). Reprogramming stem cells in regenerative medicine. *Smart Medicine*, 1(1), Article e20220005. https://doi.org/10.1002/SMMD.20220005
- Mathur, R., Thakur, K., & Hazam, R. K. (2019). Highlights of Indian council of medical research national ethical guidelines for biomedical and health research involving human participants. *Indian Journal of Pharmacology*, *51*(3), 214–221. https://doi.org/10.4103/0253-7613.262456
- Moradi, S., Mahdizadeh, H., Šarić, T., Kim, J., Harati, J., Shahsavarani, H., Greber, B., & Moore, J. B. (2019). Research and therapy with induced pluripotent stem cells (iPSCs): Social, legal, and ethical considerations. *Stem Cell Research & Therapy*, *10*(1), Article 341. https://doi.org/10.1186/s13287-019-1455-y

- Neil, H. R., Luis Gerardo Jiménez, A., & Ramón, C. (2022). Ethics of international stem cell treatments and the risk-benefit of helping patients. In D. Kitala (Ed.), *Possibilities and limitations in current Translational Stem Cell Research* (Chapter 8). IntechOpen. https://doi. org/10.5772/intechopen.108541
- Newnham, E., & Kirkham, M. (2019). Beyond autonomy: Care ethics for midwifery and the humanization of birth. *Nursing Ethics*, 26(7-8), 2147–2157. https://doi.org/10.1177/0969733018819119
- Orzechowski, M., Schochow, M., Kühl, M., & Steger, F. (2021). Content and method of information for participants in clinical studies with induced pluripotent stem cells (iPSCs). *Frontiers in cell and developmental biology*, 9, Article 627816. https://doi.org/10.3389/fcell.2021.627816
- Padilla, L. A., Hurst, D., Maxwell, K., Gawlowicz, K., Paris, W., Cleveland, D., & Cooper, D. K. (2022). Informed consent for potential recipients of pig kidney xenotransplantation in the United States. *Transplantation*, 106(9), 1754–1762. https://doi.org/10.1097/TP.0000000000004144
- Padovano, M., Scopetti, M., Manetti, F., Morena, D., Radaelli, D., D'Errico, S., Di Fazio, N., Frati, P., & Fineschi, V. (2022). Pancreatic transplant surgery and stem cell therapy: Finding the balance between therapeutic advances and ethical principles. World Journal of Stem Cells, 14(8), 577–586. https://doi.org/10.4252/wjsc.v14.i8.577
- Peng, Y.-J., Huang, X., & Zhou, Q. (2020). Ethical and policy considerations for human embryo and stem cell research in China. *Cell Stem Cell*, 27(4), 511–514. https://doi.org/https://doi.org/10.1016/j. stem.2020.09.010
- Pentaris, P., & Christodoulou, P. (2021). Qualities of culturally and religiously sensitive practice: A cross-sectional study. *Journal of palliative care*, Article 08258597211050742. https://doi.org/10.1177/08258597211050742
- Pimenta, C., Bettiol, V., Alencar-Silva, T., Franco, O. L., Pogue, R. E., Carvalho, J. L., & Felipe, M. S. S. (2021). Advanced therapies and regulatory framework in different areas of the globe: Past, present, and future. *Clinical therapeutics*, 43(5), e103–e138. https://doi.org/10.1016/j. clinthera.2021.02.006
- Resosudarmo, B. (2022). Ethics in social research in Indonesia. *Bulletin of Indonesian Economic Studies*, 58(2), 233–235. https://doi.org/10.1080/00074918.2022.2105806

- Riordan, N. H., Arias, L. G. J., & Coronado, R. (2022). Ethics of international stem cell treatments and the risk-benefit of helping patients. In D. Kitala (Ed.), *Possibilities and limitations in current translational stem cell research* (Chapter 8). IntechOpen. https://doi.org/10.5772/intechopen.108541
- Sato, K., & Suzuki, M. (2022). Standards of conducts for biostatisticians and stem cell researchers: A call for self-formulated aspirational ethics over built-in prohibitive ethics. *Science and Engineering Ethics*, *28*(2), 15. https://doi.org/10.1007/s11948-022-00366-5
- Singh, V. K., Kalsan, M., Kumar, N., Saini, A., & Chandra, R. (2015). Induced pluripotent stem cells: Applications in regenerative medicine, disease modeling, and drug discovery. *Frontiers in Cell and Developmental Biology*, 3, Article 2. https://doi.org/10.3389/fcell.2015.00002
- Søvold, L. E., Naslund, J. A., Kousoulis, A. A., Saxena, S., Qoronfleh, M. W., Grobler, C., & Münter, L. (2021). Prioritizing the mental health and well-being of healthcare workers: An urgent global public health priority. Frontiers in Public Health, 9, Article 679397. https://doi.org/10.3389/fpubh.2021.679397
- Stahl, B. C., Akintoye, S., Bitsch, L., Bringedal, B., Eke, D., Farisco, M., Grasenick, K., Guerrero, M., Knight, W., Leach, T., Nyholm, S., Ogoh, G., Rosemann, A., Salles, A., Trattnig, J., & Ulnicane, I. (2021). From responsible research and innovation to responsibility by design. *Journal of Responsible Innovation*, 8(2), 175–198. https://doi.org/10.1080/232 99460.2021.1955613
- Staunton, C., Slokenberga, S., & Mascalzoni, D. (2019). The GDPR and the research exemption: Considerations on the necessary safeguards for research biobanks. *European Journal of Human Genetics*, *27*(8), 1159–1167. https://doi.org/10.1038/s41431-019-0386-5
- Subandi, M. A., Nihayah, M., Marchira, C. R., Tyas, T., Marastuti, A., Pratiwi, R., Mediola, F., Herdiyanto, Y. K., Sari, O. K., Good, M. D., & Good, B. J. (2023). The principles of recovery-oriented mental health services: A review of the guidelines from five different countries for developing a protocol to be implemented in Yogyakarta, Indonesia. *PLoS One*, 18(3), Article e0276802. https://doi.org/10.1371/journal.pone.0276802

- Ten Have, H., & Neves, M. C. P. (2021). *Dictionary of global bioethics*. Springer. https://doi.org/10.1007/978-3-030-54161-3
- Torres-Padilla, M. E., Bredenoord, A. L., Jongsma, K. R., Lunkes, A., Marelli, L., Pinheiro, I., & Testa, G. (2020). Thinking "ethical" when designing an international, cross-disciplinary biomedical research consortium. *The EMBO Journal*, *39*(19), Article e105725. https://doi.org/10.15252/embj.2020105725
- Valdés, E., & Lecaros, J. A. (Eds.). (2023). *Handbook of bioethical decisions. Volume II: Scientific integrity and institutional ethics.* Springer Nature. https://doi.org/10.1007/978-3-031-29455-6
- van Rijssel, T. I., de Jong, A. J., Santa-Ana-Tellez, Y., Boeckhout, M., Zuidgeest, M. G., van Thiel, G. J., & Consortium, T. H. (2022). Ethics review of decentralized clinical trials (DCTs): Results of a mock ethics review. *Drug Discovery Today*, *27*(10), Article 103326. https://doi.org/10.1016/j.drudis.2022.07.011
- Varzideh, F., Gambardella, J., Kansakar, U., Jankauskas, S. S., & Santulli, G. (2023). Molecular mechanisms underlying pluripotency and self-renewal of embryonic stem cells. *International Journal of Molecular Sciences*, 24(9), Article 8386. https://doi.org/10.3390/ijms24098386
- Volarevic, V., Markovic, B. S., Gazdic, M., Volarevic, A., Jovicic, N., Arsenijevic, N., Armstrong, L., Djonov, V., Lako, M., & Stojkovic, M. (2018). Ethical and safety issues of stem cell-based therapy. *International Journal of Medical Sciences*, 15(1), 36–45. https://doi. org/10.7150/ijms.21666
- Wang, S.W., Gao, C., Zheng, Y. M., Yi, L., Lu, J. C., Huang, X. Y., Cai, J. B., Zhang, P. F., Cui, Y. H., & Ke, A. W. (2022). Current applications and future perspective of CRISPR/Cas9 gene editing in cancer. *Molecular cancer*, 21(1), Article 57. https://doi.org/10.1186/s12943-022-01518-8
- Weinbaum, C., Landree, E., Blumenthal, M. S., Piquado, T., & Gutierrez, C. I. (2019). Ethics in scientific research: An examination of ethical principles and emerging topic. RAND. https://doi.org/10.7249/RR2912
- Yamanaka, S. (2020). Pluripotent stem cell-based cell therapy: Promise and challenges. *Cell Stem Cell*, 27(4), 523–531. https://doi.org/10.1016/j. stem.2020.09.014
- Yui, H., Muto, K., Yashiro, Y., Watanabe, S., Kiya, Y., Kamisato, A., Inoue, Y., & Yamagata, Z. (2022). Comparison of the 2021 International Society for Stem Cell Research (ISSCR) guidelines for "laboratory-based human stem cell research, embryo research, and related research

- activities" and the corresponding Japanese regulations. *Regenerative Therapy*, 21, 46–51. https://doi.org/10.1016/j.reth.2022.05.002
- Żakowska-Henzler, H., Zemła-Pacud, Ż., & Zimny, T. (2023). *Biotechnology, patents and human rights in Europe: Innovations concerning the human body*. Edward Elgar Publishing. https://doi.org/10.4337/9781803920269
- Zalaf, B. R., Bringel, M., Jorge, P. K., de Oliveira, B., Tanabe, K., Santos, C. F., Oliveira, R. C. d., Rios, D., Cruvinel, T., Lourenço Neto, N., Oliveira, T. M. d., & Machado, M. A. d. A. M. (2020). A biobank of stem cells of human exfoliated deciduous teeth: Overview of applications and developments in Brazil. *Cells Tissues Organs*, 209, 37–42. https://doi.org/10.1159/000506677
- Zarzeczny, A. (2019). The future of stem cell research and its clinical translation in Canada: Exploring questions of governance and policy options. In K. Turksen (Ed.), *Advances in experimental medicine and biology* (1–16). Springer.