



A SYSTEMATIC REVIEW OF DEVELOPMENTAL OUTCOMES IN CHILDREN WITH DOWN SYNDROME.

Wenuri Thamalka Kodagoda

Faculty of Nursing

Australian College of Business and Technology

Sri Lanka.

Abstract

Background: Down syndrome is a genetic condition that affects how children develop in terms of their thinking, health, and social skills. While early help, inclusive education, and family support have been shown to improve development, there are still gaps in understanding the long-term effects of health problems and how different environments affect children. Additionally, there's not enough research from regions like South Asia, Africa, and Latin America, where children might face different challenges.

Purpose and Method: This systematic review looks at 53 studies published between 1996 and 2024 to understand how children with Down syndrome develop. The study uses bibliometric methods to find trends, important researchers, and areas where more research is needed. The focus is on how cognitive, health, and social factors affect development. This abstract also examines the role of culturally specific interventions in improving developmental outcomes in diverse regions.

Results: The review shows that early interventions, inclusive education, and family support help improve cognitive skills, language, problem-solving, and social skills. However, there are still many gaps in understanding how health issues, like sleep disorders and oxidative stress, affect learning over time. The review also highlights that research is lacking from underrepresented regions. New technologies, such as apps and digital tools, are being explored to help children with Down syndrome improve communication and social skills, especially in areas with fewer resources.

Value: This review highlights the importance of a team approach, including healthcare providers, teachers, and families, in supporting children with Down syndrome. The study emphasizes the need for standardization in measuring outcomes to enable global comparisons across studies. The findings suggest that addressing the current gaps in research and using new technologies can help these children develop important life skills and live more independent and fulfilling lives. More research is needed to explore long-term solutions, especially in underrepresented regions and areas like health and social development.

Index Terms: Down syndrome, developmental outcomes, early intervention, education, family support, health challenges, social development.

INTRODUCTION

This review seeks to analyze the developmental outcomes in children with Down syndrome using bibliometric methods.

Down syndrome is a genetic condition that occurs when a child has an extra copy of chromosome 21. This extra chromosome leads to a range of developmental challenges affecting cognitive abilities, physical health, and social skills. Children with Down syndrome may experience delays in language development, motor

skills, and may face cognitive difficulties, but the severity of these challenges can vary significantly among individuals. Over the years, extensive research has been conducted to understand the factors that influence developmental outcomes in children with Down syndrome, focusing on the impact of early intervention, healthcare, education, and social support systems. However, despite the progress made, it remains unclear why some children with Down syndrome achieve advanced cognitive and social skills, while others experience more severe delays (Lukowski et al., 2019).

This difference in outcomes highlights the complexity of Down syndrome and suggests that further research is needed to uncover the specific factors that contribute to these varying results. Understanding how cognitive, health, and environmental factors interact and affect developmental progress is crucial for improving the quality of life for children with Down syndrome. Early intervention programs, inclusive educational practices, and strong family support have been shown to improve developmental outcomes, but more work is needed to identify which factors have the most significant impact and how they work together (Grane et al., 2023).

The developmental outcomes of children with Down syndrome are shaped by a combination of genetic, health, cognitive, and environmental factors. While some children show strong progress in areas like cognitive function, language, and social development, others face significant barriers that affect their ability to engage in daily activities and learn effectively. Research has shown that early medical interventions and therapies, including physical, occupational, and speech therapies, can have a positive impact on a child's developmental progress. However, the long-term effects of health issues such as sleep disorders, heart problems, and oxidative stress on cognitive development have not been sufficiently explored (Chawla et al., 2020).

Another key factor that influences developmental outcomes is the child's environment, particularly their access to quality healthcare, educational opportunities, and emotional support from their families and communities. Studies have suggested that inclusive education—where children with Down syndrome learn alongside their typically developing peers—can help improve social and cognitive skills. However, there is a lack of research on how family support and community involvement affect developmental outcomes over the long term. Many studies focus on one specific aspect of development, whether it be cognitive skills or health-related issues, but fail to examine how these factors work together in a child's overall development. This introduction also highlights the underrepresentation of studies in low-income regions, which face distinct challenges due to limited resources and cultural differences.

This review seeks to synthesize existing literature and provide a more integrated approach to understanding the developmental outcomes in children with Down syndrome. By examining cognitive, health, and environmental factors together, this review aims to offer a comprehensive view of how these elements influence a child's development and identify areas where more focused research is needed to improve outcomes for children with Down syndrome. It emphasizes the growing need for cross-disciplinary research combining genetics, environmental science, and behavioral studies to understand the multifaceted nature of Down syndrome.

PROBLEM STATEMENT

Down syndrome is a genetic condition that significantly impacts children's cognitive, physical, and social development. While various interventions and support systems have been implemented to improve developmental outcomes, their effectiveness varies widely among children. Factors such as individual health conditions, access to healthcare and education, and family or community support can influence these outcomes. Despite ongoing research, there is limited understanding of how these factors interact and contribute to developmental progress in children with Down syndrome. This study aims to examine the influences of cognitive, health, and environmental factors on developmental outcomes, identifying strategies to better support children with Down syndrome and enhance their quality of life.

RESEARCH QUESTIONS:

1. What are the main trends, themes and geographical patterns in the research on developmental outcomes in children with down syndrome?
2. What are the research gap and the limitations?
3. Who are the most influential authors and impactful journals?
4. What are the key words occurrences in the research on developmental outcomes in children with down syndrome?
5. What are the key authors occurrences in the research on developmental outcomes in children with down syndrome?

RESEARCH OBJECTIVES:

1. To identify the main trends, themes and geographical patterns in the research on developmental outcomes in children with down syndrome.
 2. To examine the research gap and the limitations.
 3. To find the most influential authors and impactful journals.
 4. To analysis the key words occurrences in the research on developmental outcomes in children with down syndrome.
- To determine the key authors occurrences in the research on developmental outcomes in children with down syndrome.

METHODOLOGY

This systematic review analyzed 102 research articles published between 1996 and 2024 to explore developmental outcomes in children with Down syndrome (Bunt & Bunt, 2014). The articles came from several academic sources, including PubMed, Emerald Insight, Semantic Scholar, DOI.org (CrossRef), MDPI, Wiley Online Library, SciELO, and ResearchGate. From these, PubMed provided 37 articles, Emerald Insight 17 articles, Semantic Scholar 18 articles, DOI.org 12 articles, MDPI 2 articles, and Wiley Online Library 8 articles, SciELO 4 articles, and ResearchGate 4 articles each. Ultimately, 53 papers were reviewed for this research.

Phase 1 – Data Mapping through Keyword Search

The first step in the process involved identifying the relevant studies using key terms. A combination of keywords such as "Down syndrome," "developmental outcomes," "cognitive development," "health challenges," and "environmental factors" were used. Boolean operators (AND, OR) were applied to refine the search and ensure more accurate results. Filters were set to limit the search to studies published in English and focusing on children aged 0–18 years, which is the critical developmental period for children with Down syndrome. Zotero software was used to organize and manage all the references collected during this phase. This allowed for efficient tracking of articles and ensured proper citation handling throughout the process. Additionally, various databases were selected to provide a comprehensive range of perspectives on the topic, allowing for a broad analysis of the literature. The combination of these resources provided a more well-rounded view of the developmental outcomes in children with Down syndrome.

Phase 2 – Refining Results

In this phase, studies were selected based on specific inclusion and exclusion criteria. Only those studies that focused on the impact of cognitive, health, or environmental factors on children with Down syndrome were included. Cognitive studies examined developmental areas such as memory, language acquisition, and problem-solving. Health studies investigated concerns like sleep problems, metabolic issues, and oxidative stress (Esbensen et al., 2016). Environmental studies addressed factors such as the role of family support and access to educational opportunities. Studies that focused on adults, non-peer-reviewed articles, or those that were not fully available in text were excluded from the review. After applying these criteria, a total of 53 studies were retained for further analysis. This filtering process ensured that only the most relevant and high-quality studies were included, contributing to the reliability of the review's findings. This process also helped ensure that the research outcomes were as accurate as possible, providing a clear picture of the developmental challenges children with Down syndrome face.

Phase 3 – Article Assessment and Review Conclusions

In this final phase, the quality and reliability of the selected studies were evaluated using the Joanna Briggs Institute checklist. This tool helped assess the methodological quality of the studies, ensuring that only rigorous and reliable research was included. The selected articles were then categorized into three primary areas: cognitive development, health outcomes, and environmental factors (Baxter et al., 2022). Bibliometric Analysis, VOSviewer software was used for bibliometric analysis, providing insights into key authors, impactful journals, and recurring research themes.

Additionally, the following analyses were included:

- 1.A PRISMA Chart illustrated the inclusion and exclusion process.
- 2.A Geographical Distribution Map showed research activity by region, highlighting underrepresented areas like South Asia and Africa (Aggeliki, 2019).
- 3.A Growth Trend Chart depicted an increase in research from 1996 to 2024, with notable growth after 2014.
- 4.A Methodological Breakdown Chart displayed the proportions of quantitative (54%), qualitative (24%), and mixed methods (22%) studies (Baxter et al., 2022).

5.A Keyword Co-occurrence Map visualized frequently discussed terms such as "development," "child," and "treatment," using VOSviewer (Chawla et al., 2020).

6.A Co-Authorship Network Chart revealed global collaboration patterns among researchers (Hanna et al., 2022).

7.A Publication Source Chart highlighted databases, with PubMed contributing the most studies (Esbensen et al., 2016).

8.Citations Analysis: A detailed review of the most influential authors and journals based on citation counts (Grane et al., 2023).

9.University Distribution Analysis: A map highlighted global collaborations, including key institutions like Monash University, University of Pennsylvania, and Sapienza University of Rome, reflecting diverse research contributions (Van Riper & Cohen, 2001).

PRISMA FLOWCHART.

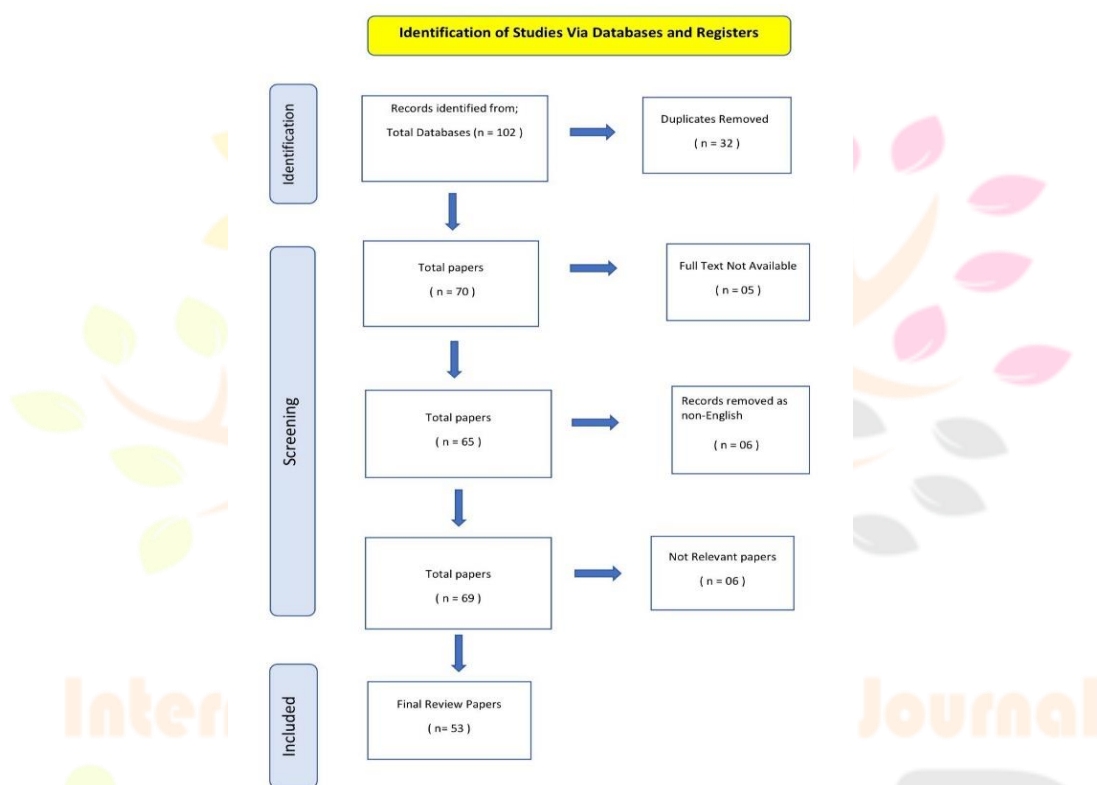


Figure 1: PRISMA Chart

GROWTH OF RESEARCH INTEREST OVER THE YEARS

The following figure (Figure 2) illustrates the growth of research on developmental outcomes in children with Down syndrome from 1996 to 2024. The chart shows that early years saw minimal publications, with only one or two articles published annually. However, starting around 2014, there was a noticeable increase in research activity, reaching a peak in 2024 with 9 publications.

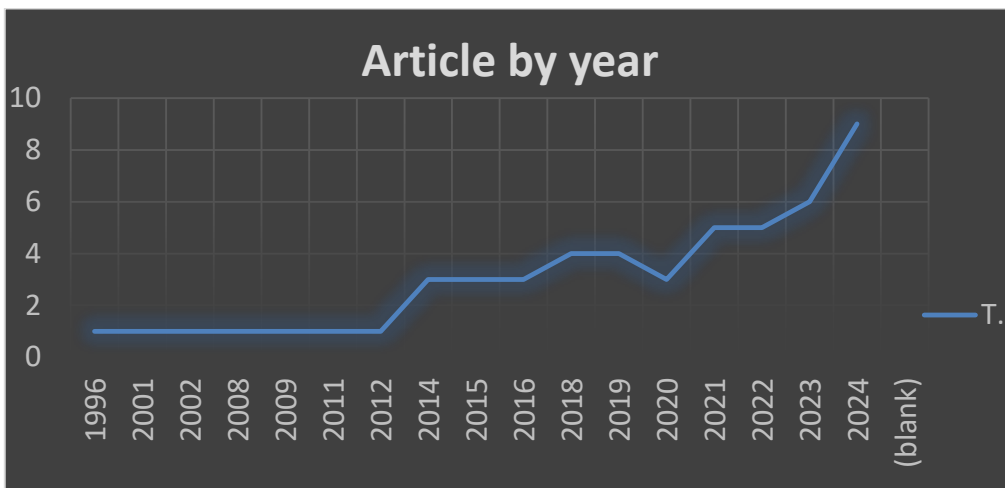


Figure 2: Growth of research interest on developmental outcomes in children with Down Syndrome (1996–2024).

This increase in research publications highlights a growing focus on understanding the developmental challenges faced by children with Down syndrome. The rise in research activity, particularly after 2014, may be due to the heightened awareness of the need for better interventions and support for these children. It reflects the expanding body of knowledge surrounding their cognitive, health, and environmental outcomes, which have become central areas of study in recent years. The surge in publications in 2024 suggests that the research community is now dedicating more resources and attention to this important topic, signaling an ongoing trend of deepening interest in the field.

NUMBER OF CITATIONS BY PAPER TYPE OVER THE YEARS

Figure 3 shows the number of studies by type (Quantitative, Qualitative, and Mixed) across different years. There are 50 studies in total: 27 are quantitative, 12 are qualitative, and 11 are mixed. The year 2024 has the highest number of studies, with 9 contributions (6 quantitative, 1 qualitative, and 2 mixed). In earlier years like 2008, 2002, 2001, and 1996, only 1 study was published each year.

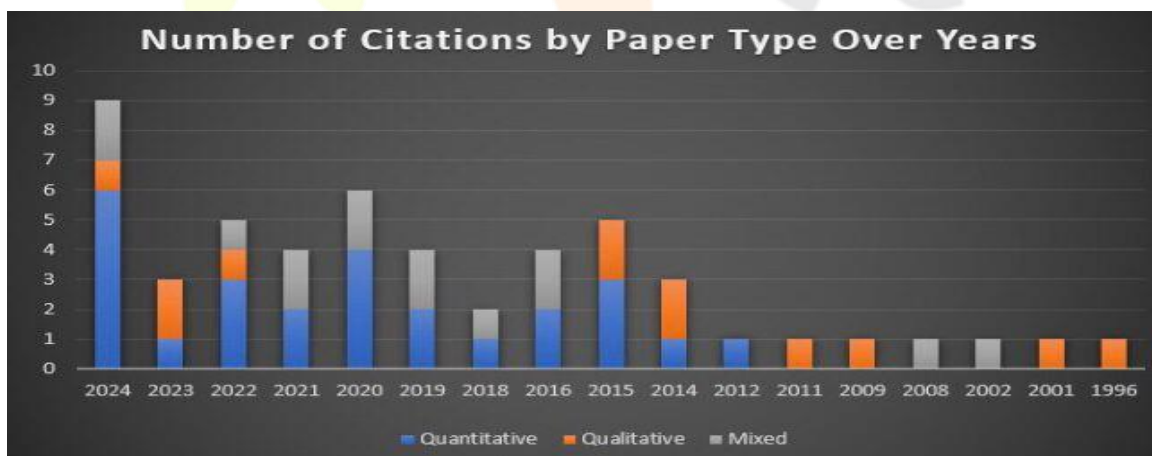


Figure 3: Number of Citations by Paper Type Over the Years.

The chart highlights a growing trend in quantitative and mixed-method studies in recent years. It shows how research methods have evolved over time, with a clear focus on quantitative studies throughout most years. This makes it easy to see patterns and changes in research activity.

PUBLISHED SITES

The following figure (Figure 4) shows the number of publications from different sites used by authors. Most publications are from PubMed, with 36 entries, showing its key role in medical and scientific research. Semantic Scholar and www.emerald.com each contribute 6 entries, highlighting their importance in multidisciplinary studies. Other sources include Wiley Online Library and SciELO, with 1 entry each, and www.mdpi.com, with 2 entries.

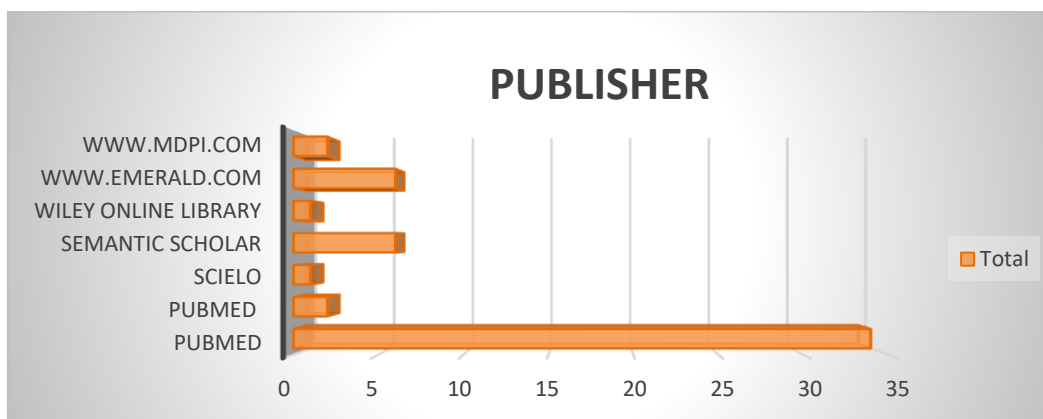


Figure 4: Bar chart of Publisher.

This chart highlights the heavy reliance on PubMed, while also using other platforms like Semantic Scholar and MDPI to support research. It shows a diverse approach to gathering information from both specialized and general academic resources.

CREDIBILITY OF JOURNALS.

This research, which systematically reviews developmental outcomes in children with Down syndrome, highlights the importance of using reliable and high-quality studies from trusted sources. These studies cover key areas such as cognitive, social, and health-related development. By selecting credible journals, the findings are supported by evidence that has undergone thorough review. Measures like the H-index, Impact Score, and SJR demonstrate the quality and academic influence of the journals, as well as the impact of the research they publish.

The table below presents the selected journals for this review, along with their respective H-index values and other relevant metrics, to showcase their credibility and significance in developmental research.

<u>Jounarl</u>	No: Articles	Impact Score	H-Index	SJR	Overall Ranking
Journal of Clinical Medicine	1	0.78	14	0.222	18829
<u>Revista da Associação Médica Brasileira</u>	1	0.81	47	0.314	15323
Singapore Medical Journal	1	0.89	71	0.374	13678
Journal of Enabling Technologies	1	1.86	24	0.394	13133
Advances in Mental Health and Intellectual Disabilities	1	1.50	18	0.433	12200
<u>Roczniki Panstwowego Zakladu Higieny</u>	1	1.39	28	0.29	16033
Journal of Cognitive Psychology	1	1.65	73	0.507	10630
Nursing Children and Young People	1	0.58	25	0.192	20411
American Family Physician	1	1.09	148	0.546	9904

METHODOLOGICAL ANALYSIS.

The research methods used in studies about children with Down syndrome include three types: Quantitative, Qualitative, and Mixed methods. Most studies (54%) used Quantitative methods, which focus on numbers and data to measure areas like thinking skills and health issues. These methods are helpful for providing clear results but don't show much about personal or family experiences.

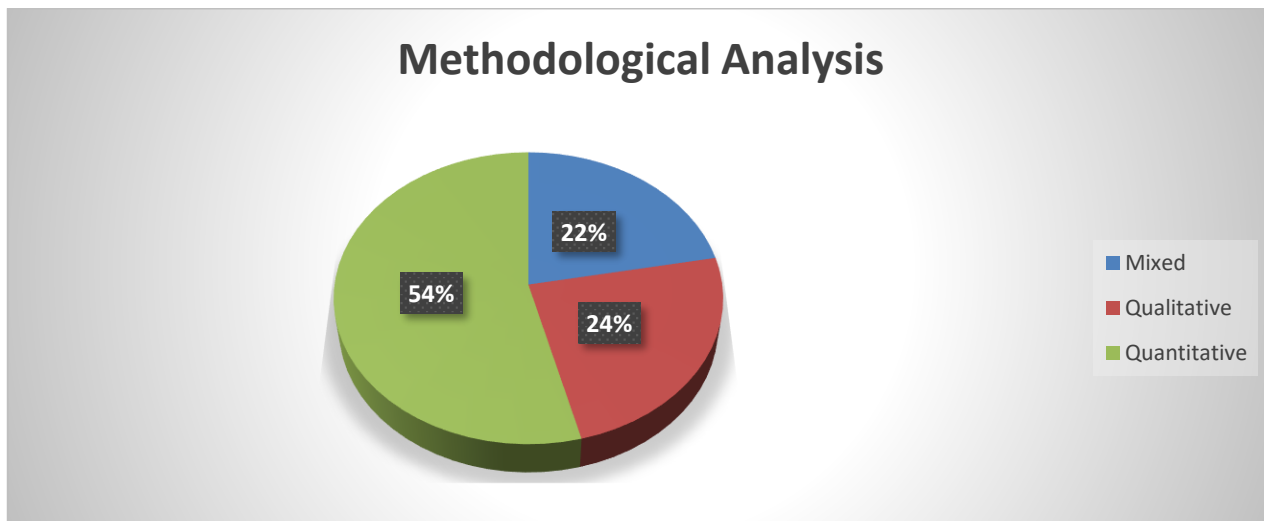


Figure 5: Methodological Analysis.

Some studies (24%) used Qualitative methods, which explore experiences and feelings, such as how families or schools support children. Only 22% used Mixed methods, which combine both quantitative and qualitative approaches. Mixed methods give a fuller understanding of how health, thinking, and family factors connect, but they are not used enough. Future research should focus more on mixed methods to provide better insights.

MOST INFLUENTIAL AUTHORS AND CITATIONS

A detailed study was done to find the most influential authors in this area of research. From 53 articles, over 120 authors were identified for their contributions. Some authors stood out because their work was cited more often, showing their strong impact and influence. The table below highlights the top authors and their citation counts, showing how important they are in advancing this field.

1	AUTHORS	CITATIONS
2	Asim	490
3	Zemel	249
4	Coppedè	168
5	Agarwal Gupta	117
6	Næss	111
7	Lee	89
8	Nordstrøm	72
9	Lukowski	62
10	Rosso	53
11	Van Riper	51

COUNTRY OF ARTICLE PUBLICATIONS

The following figure (Figure 6) illustrates the distribution of research on developmental outcomes in children with Down syndrome across different countries. The chart shows that the research landscape is quite diverse, with contributions from a range of countries worldwide. Some countries, like the United States and the United Kingdom, have notably higher publication rates, while others have fewer contributions.



Figure 6: Distribution of article publications by country.

The United States leads with the highest number of publications (10), followed by the United Kingdom (4), and Australia (4). This highlights that English-speaking nations are at the forefront of research in this area. However, countries like India, Brazil, and Japan also show significant contributions, reflecting the international interest in understanding the developmental outcomes of children with Down syndrome. This diverse geographical representation underscores the global significance of this research.

DISTRIBUTION OF UNIVERSITIES REPRESENTED BY THE AUTHORS.

The dataset includes 53 entries highlighting the university's global collaborations with countries such as Australia, Turkey, the United States, Germany, Italy, Norway, Japan, Saudi Arabia, Greece, Portugal, and the UK. These partnerships focus on important areas like mental health, child development, pediatrics, and neuropsychology. By working across regions like Europe, Asia, North America, and Africa, the university fosters knowledge sharing and cultural exchange to address global challenges.



Figure 7: Distribution of Universities represented by the Authors.

Key partner universities include Monash University, Torrens University, and Western Sydney University in Australia; Samsun University in Turkey; Michigan State University, the University of Cincinnati, and the University of Pennsylvania in the United States; and Radboud University Medical Center in the Netherlands. Collaborations also include the University of Hertfordshire (UK), Yamaguchi University (Japan), Sapienza University of Rome and Bambino Gesù Children's Hospital (Italy), King Saud University (Saudi Arabia), and the University of Coimbra (Portugal), among others.

CO- AUTHOR OCCURRENCE ANALYSIS

To understand how researchers work together on the topic “A Systematic Review of Developmental Outcomes in Children with Down Syndrome,” a co-authorship network was created using the VOS viewer tool. In Figure 8, each circle shows an author, and the size represents how many papers they wrote or how much they collaborated with others. The lines between the circles show connections, meaning those authors worked together. The colors group authors who often collaborate, forming research teams.

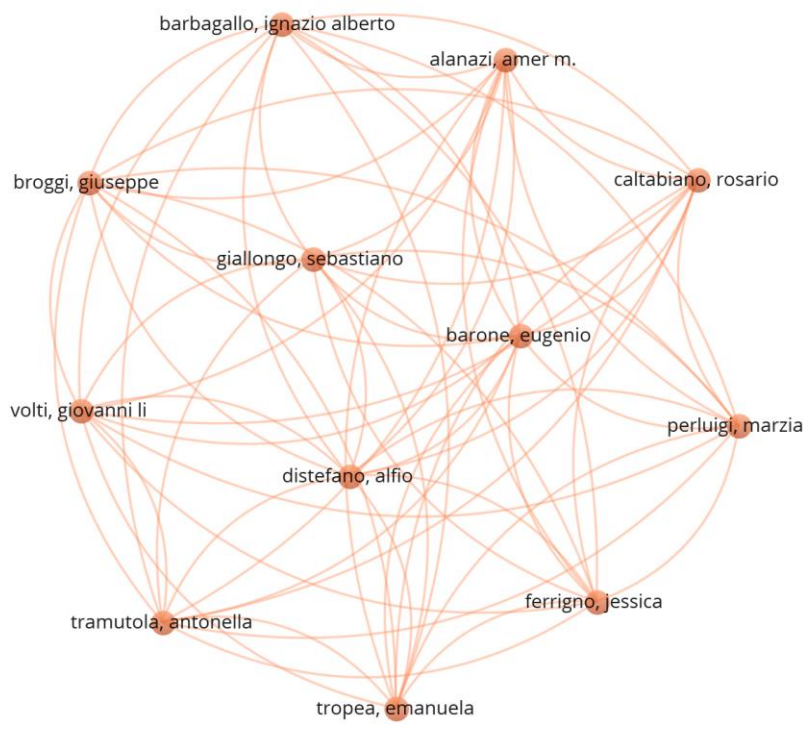
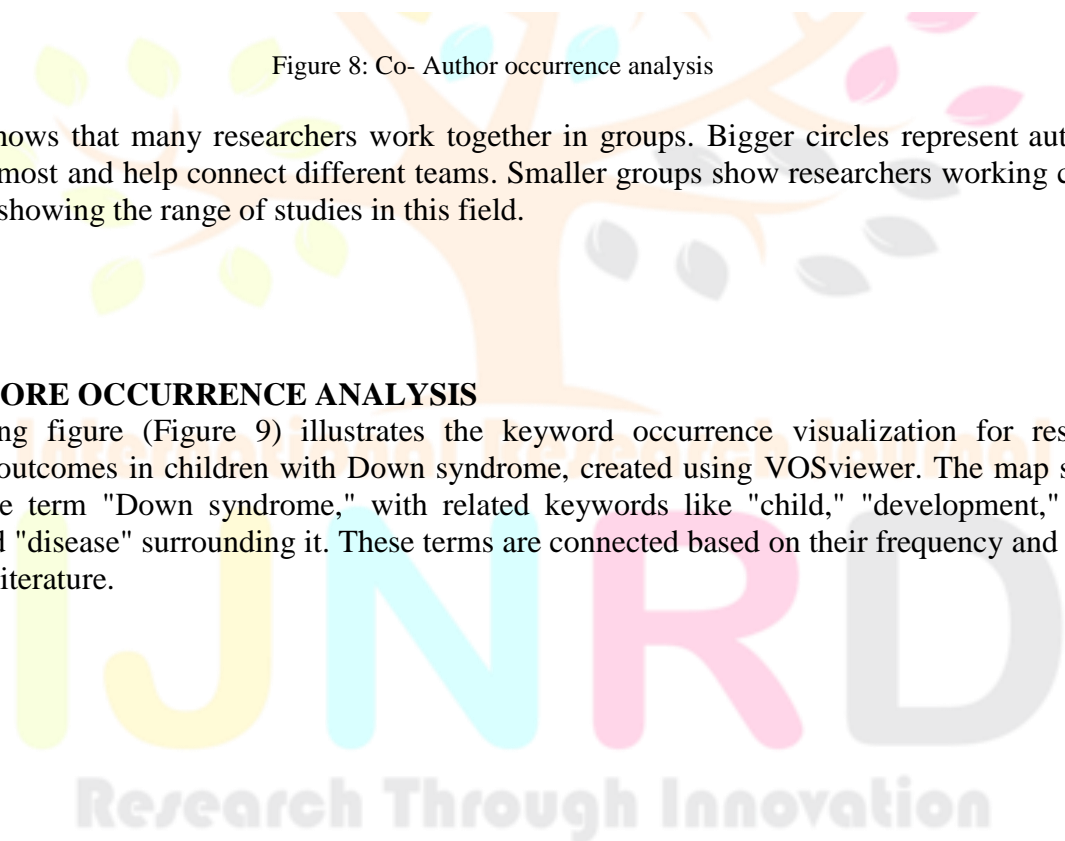


Figure 8: Co- Author occurrence analysis

This chart shows that many researchers work together in groups. Bigger circles represent authors who collaborate the most and help connect different teams. Smaller groups show researchers working closely on specific topics, showing the range of studies in this field.

KEYWORD CORE OCCURRENCE ANALYSIS

The following figure (Figure 9) illustrates the keyword occurrence visualization for research on developmental outcomes in children with Down syndrome, created using VOSviewer. The map shows the centrality of the term "Down syndrome," with related keywords like "child," "development," "parent," "treatment," and "disease" surrounding it. These terms are connected based on their frequency and relevance in the research literature.



RESEARCH GAPS & LIMITATIONS.

Research Gaps.

1. Knowledge Gaps

- We don't fully understand why some children with Down syndrome develop better thinking and learning skills than others.
- There is little research on how health problems like sleep-disordered breathing and oxidative stress impact cognitive development over time.
- There's not enough research on how thinking skills, health, and family support work together.

2. Methodological Gaps

- Many studies are too small, so the results don't apply to all children with Down syndrome.
- Different studies use different ways to measure development, so it's hard to compare them.
- Researchers don't often combine numbers and personal stories, even though it could give better answers.

3. Geographical Gaps

- Most studies are done in rich countries like the US and UK. Few come from places like South Asia, Africa, or Latin America, where children may face different challenges.

Limitations.

- Focus on singular factors rather than integrated analysis.
- Limited access to comprehensive datasets from low-resource regions.
- Insufficient evaluation of innovative tools like digital therapies in diverse contexts.

DISCUSSION

This review analyzed 53 studies to understand how children with Down syndrome grow and develop. It focused on four key factors: cognitive development, health issues, environmental influences, and family support.

1. Cognitive Development

Children with Down syndrome show many differences in how they learn and think. Some children do well with problem-solving and language, while others find these areas harder (Lukowski et al., 2019). Early interventions, such as speech therapy and inclusive education, can help improve these skills. Inclusive classrooms allow children with Down syndrome to learn alongside their peers, which helps them build confidence and social skills.

Using technology, like apps and online learning tools, is another way to support cognitive development. For example, apps designed to teach language or improve memory can be very helpful (Prena & Sherry, 2018). These tools are especially useful in areas where access to therapists or special programs is limited. However, more research is needed to understand how well these technologies work in different parts of the world.

2. Health Issues

Health problems are common for children with Down syndrome and can affect their learning and growth. Sleep disorders, such as sleep apnea, can make children feel tired and affect their ability to focus and remember things (Hanna et al., 2022). Treating sleep problems early can lead to better learning outcomes.

Oxidative stress is another issue that can slow down development. It is linked to behavior problems and learning delays (Chawla et al., 2020). Proper medical care, regular check-ups, and a healthy diet can help reduce these effects. Nutritional problems, like obesity or poor diet, are also common and can make health problems worse. Helping families learn about good nutrition and healthy habits can improve both physical and mental development (Nordstrøm et al., 2020).

3. Environmental Influences

The environment a child grows up in plays a big role in their development. Children with access to good healthcare, inclusive schools, and safe communities tend to do better. However, many children in regions like South Asia, Africa, and Latin America do not have these opportunities (Aggeliki, 2019). This lack of resources makes it harder for them to reach their potential.

Governments and organizations need to focus on creating more inclusive schools and making healthcare affordable. Community programs can also teach families about the services available to them, helping them make better choices for their children.

4. Family Support

Families are one of the most important influences on a child's growth. A supportive family helps children feel secure and motivated to learn. Parents who are involved in their child's therapy and education often see better results (Van Riper & Cohen, 2001).

However, families can face challenges, especially in areas with fewer resources. Parents may feel stressed or overwhelmed when they cannot find the right support for their child. Programs that provide counseling, financial help, and parent training can make a big difference. Families need to feel supported so they can provide the best care for their children.

By addressing these four factors—cognitive development, health, environment, and family support—we can help children with Down syndrome grow and thrive. This will require global efforts, new technologies, and programs designed to meet the needs of children and families in different parts of the world.

FUTURE RESEARCH DIRECTIONS

1. Health and Development: Researchers should study how health issues, like poor sleep, impact learning and behavior. Understanding these links can help develop better treatments for children with Down syndrome.

2. Underrepresented Regions: There isn't much research from areas like South Asia, Africa, and Latin America. Studies in these regions are needed to understand the unique challenges faced by children in different cultures and environments.

3. Comprehensive Programs: Future research should test programs that combine health treatments, education, and family support. This combined approach could be more effective than focusing on one area alone.

4. Use of Technology: Digital tools, such as apps and remote support programs, could help children with Down syndrome, especially in areas with fewer resources. Researchers should explore how technology can improve access to healthcare and education.

CONCLUSION.

This review shows that the development of children with Down syndrome depends on a mix of cognitive, health, and environmental factors. While progress has been made, there are still gaps in understanding how these factors work together (Baumer & Capone, 2023). Access to good healthcare, education, and family support is essential for positive outcomes. However, many children, especially in poorer regions, don't have access to these resources (Marqui & Borges, 2024). More studies are needed in these areas to improve support for children with Down syndrome worldwide.

Future research should focus on programs that combine health, education, and family care. Technology should also be used to improve access to services (Souza de Oliveira et al., 2024). By addressing these gaps, we can help children with Down syndrome live healthier and happier lives.

It is also important to explore how early interventions can better support children's cognitive development. Programs like speech therapy, occupational therapy, and inclusive education have shown to improve language and problem-solving skills significantly (Lukowski et al., 2019). More research is needed to understand how these programs can be adapted for children in different cultural and economic settings (Aggeliki, 2019).

Health challenges such as sleep disorders and oxidative stress also require more attention. These issues are linked to learning delays and behavior problems (Chawla et al., 2020). Addressing these health problems early can lead to better cognitive and social development outcomes. For example, improving sleep quality through treatments or therapies can enhance focus and memory in children with Down syndrome (Hanna et al., 2022).

Finally, researchers should study the role of family support more deeply. A strong family environment, combined with access to community resources, helps children adapt better socially and emotionally (Van Riper & Cohen, 2001). Studies in regions like South Asia and Africa are especially important to understand how local factors affect children's development (Grane et al., 2023).

By combining health, education, and family-focused solutions, we can create better opportunities for children with Down syndrome. This effort will require global cooperation and ongoing research, but it is key to helping these children lead independent and fulfilling lives.

REFERENCES

- Agarwal Gupta, Neerja, and Madhulika Kabra. 2014. "Diagnosis and Management of Down Syndrome." *Indian Journal of Pediatrics*, Down syndrome, diagnosis, management, health interventions, intellectual disability, 81 (6): 560–67. <https://doi.org/10.1007/s12098-013-1249-7>.
- Aggeliki, Sideraki. 2019. "Down Syndrome, Characteristics, Diagnosis, Education, and the Role of the Family." *The Study Focuses on Children with Down Syndrome, Their Families, and Educational Practitioners Involved in Their Care.*, Down syndrome, characteristics, diagnosis, education, family role, genetic factors, developmental stages, , October, 69–77.
- Akça, Gülfer, Aslihan Sanri, and Unal Akca. 2024. "Health Literacy in Parents of Children with Down Syndrome." *Advances in Mental Health and Intellectual Disabilities*, Down syndrome, health literacy, genetic counseling, parent education., 18 (2): 88–97. <https://doi.org/10.1108/AMHID-10-2023-0038>.
- Alshammar, Abdullah Khalaf A, Salma Sameer A Alkattan, Rahaf Mohammed S Alsharif, Nawaf Falah J Alwahbi, Kawther Ali A Alhussain, Ahmad Mohammed M Alqahtani, Abdulmajeed Hussain Saedi, Mohammed Abdullah M Alhussain, Mohammad Hassan Haroobi, and Muath Ali H Alshehri. 2021. "Down Syndrome Clinical Features, and It's Associated Complications Evaluation and Management Approach." *Pharmacophore*, Down syndrome, clinical features, complications, management, evaluation, genetic disorder, 12 (4): 103–6. <https://doi.org/10.51847/n81OfVwvph>.
- Anil, M. A., S. Shabnam, and S. Narayanan. 2019. "Feeding and Swallowing Difficulties in Children with Down Syndrome." *Journal of Intellectual Disability Research: JIDR*, Feeding difficulties, swallowing, Down syndrome, oral motor skills, dysphagia, intervention., 63 (8): 992–1014. <https://doi.org/10.1111/jir.12617>.
- Aoki, Sayaka, Yuko Yamauchi, and Keiji Hashimoto. 2018. "Developmental Trend of Children with Down's Syndrome - How Do Sex and Neonatal Conditions Influence Their Developmental Patterns?" *Brain & Development*, : Down syndrome, developmental trends, sex differences, neonatal conditions, cognitive development., 40 (3): 181–87. <https://doi.org/10.1016/j.braindev.2017.10.001>.
- Asim, Ambreen, Ashok Kumar, Srinivasan Muthuswamy, Shalu Jain, and Sarita Agarwal. 2015. "Down Syndrome: An Insight of the Disease." *Journal of Biomedical Science*, Down syndrome, trisomy 21, genetic disorders, congenital defects, cardiac problems, leukemia, hypertension, gastrointestinal issues, genetic mutations., 22 (1): 41. <https://doi.org/10.1186/s12929-015-0138-y>.
- Baumer, Nicole T., and George Capone. 2023. "Psychopharmacological Treatments in Down Syndrome and Autism Spectrum Disorder: State of the Research and Practical Considerations." *American Journal of Medical Genetics. Part C, Seminars in Medical Genetics*, Psychopharmacological treatments, Down syndrome, autism spectrum disorder, behavioral symptoms, clinical trials, psychiatric symptoms., 193 (4): e32069. <https://doi.org/10.1002/ajmg.c.32069>.
- Baxter, Rebecca, Rachel Rees, Alexandra Perovic, and Charles Hulme. 2022. "The Nature and Causes of Children's Grammatical Difficulties: Evidence from an Intervention to Improve Past Tense Marking in Children with Down Syndrome." *Developmental Science*, Down syndrome, past tense marking, language difficulties, grammar intervention, children, language acquisition., 25 (4): e13220. <https://doi.org/10.1111/desc.13220>.
- Bochud-Fragnière, Emilie, Paola Bittolo, Giada Ehrensperger, Nicole Antonicelli, Floriana Costanzo, Deny Menghini, Stefano Vicari, Pamela Banta Lavenex, and Pierre Lavenex. 2024. "Conditional Learning Abilities in Down Syndrome and Williams Syndrome." *Journal of Cognitive Psychology*, Conditional learning, Down syndrome, Williams syndrome, cognitive development, neuropsychological testing, hippocampus, 36 (7): 844–66. <https://doi.org/10.1080/20445911.2024.2390538>.
- Booster, Genery, Stephanie Jump, and Lisa Meltzer. 2022. "0516 Characterizing Sleep and Mood during COVID for Youth with Allergic Disease." *Sleep*, Sleep, mood, COVID-19, allergic diseases, youth, asthma, eczema., 45 (Supplement_1): A228–A228. <https://doi.org/10.1093/sleep/zsac079.513>.
- Boylu, Muhammed Emin, İlker Taşdemir, Mehmet Doğan, and Alaattin Duran. 2024. "What Is Important in Forensic Psychiatric Evaluation in People with Down Syndrome? A Sample from Türkiye." *Journal of Intellectual Disabilities and Offending Behaviour*, Forensic psychiatry, Down

syndrome, psychiatric evaluation, legal settings, cognitive impairments, 15 (1/2): 1–13. <https://doi.org/10.1108/JIDOB-11-2023-0008>.

- Bunt, Christopher W., and Stephanie K. Bunt. 2014. “Role of the Family Physician in the Care of Children with Down Syndrome.” *American Family Physician*, Family physician, Down syndrome, health supervision, early intervention, screenings, developmental milestones, 90 (12): 851–58.
- Chawla, Jasneek K., Anne Bernard, Helen Heussler, and Scott Burgess. 2021. “Sleep, Function, Behaviour and Cognition in a Cohort of Children with Down Syndrome.” *Brain Sciences*, Down syndrome, sleep disorders, obstructive sleep apnea, behavioral outcomes, cognitive outcomes, child sleep habits, non-respiratory sleep problems., 11 (10): 1317. <https://doi.org/10.3390/brainsci11101317>.
- Chawla, Jasneek K., Scott Burgess, and Helen Heussler. 2020. “The Impact of Sleep Problems on Functional and Cognitive Outcomes in Children with Down Syndrome: A Review of the Literature.” *Journal of Clinical Sleep Medicine: JCSM: Official Publication of the American Academy of Sleep Medicine*, Sleep disorders, cognition, behavior, Down syndrome, functional status, childhood diseases, sleep patterns, systematic review, 16 (10): 1785–95. <https://doi.org/10.5664/jcsm.8630>.
- Coppedè, Fabio. 2016. “Risk Factors for Down Syndrome.” *Archives of Toxicology*, Down syndrome, risk factors, maternal age, genetics, translocation, folate metabolism, chromosome 21., 90 (12): 2917–29. <https://doi.org/10.1007/s00204-016-1843-3>.
- Crawford, Doreen, and Annette Dearnun. 2016. “Down’s Syndrome.” *Nursing Children and Young People* 28 (9): 17. <https://doi.org/10.7748/ncyp.28.9.17.s19>.
- Dotan, Miri, Elena Zion, Haim Ben-Zvi, Havatzelet Yarden-Bilavsky, and Efraim Bilavsky. 2021. “Adenovirus Infection in Children with Down Syndrome.” *The Israel Medical Association Journal: IMAJ* 23 (7): 416–19.
- Downes, Alison, Julia S. Anixt, Anna J. Esbensen, Susan Wiley, and Jareen Meinen-Derr. 2015. “Psychotropic Medication Use in Children and Adolescents With Down Syndrome.” *Journal of Developmental and Behavioral Pediatrics: JDBP*, psychotropic medication, Down syndrome, children, adolescents, mental health, developmental disabilities, 36 (8): 613–19. <https://doi.org/10.1097/DBP.0000000000000179>.
- Esbensen, Anna J., Dean W. Beebe, Kelly C. Byars, and Emily K. Hoffman. 2016. “Use of Sleep Evaluations and Treatments in Children with Down Syndrome.” *Journal of Developmental and Behavioral Pediatrics: JDBP*, Sleep problems, Down syndrome, Polysomnography (PSG), Sleep treatments, Obstructive sleep apnea, Behavioral sleep therapy, Medical intervention., 37 (8): 629–36. <https://doi.org/10.1097/DBP.0000000000000333>.
- Giallongo, Sebastiano, Jessica Ferrigno, Rosario Caltabiano, Giuseppe Broggi, Amer M. Alanazi, Alfio Distefano, Emanuela Tropea, et al. 2024. “Aging Exacerbates Oxidative Stress and Liver Fibrosis in an Animal Model of Down Syndrome.” *Aging (Albany NY)*, Down Syndrome, oxidative stress, liver fibrosis, aging, lipid metabolism., 16 (12): 10203–15. <https://doi.org/10.18632/aging.205970>.
- Grane, Fiona Mc, Fiona Lynn, Joanne Balfe, Eleanor Molloy, and Lynne Marsh. 2023. “Down Syndrome: Parental Experiences of a Postnatal Diagnosis.” *Journal of Intellectual Disabilities: JOID*, Down syndrome, postnatal diagnosis, parents, emotional responses, healthcare delivery, 27 (4): 1032–44. <https://doi.org/10.1177/17446295221106151>.
- Hanna, Nardin, Youstina Hanna, Henrietta Blinder, Julia Bokhaut, and Sherri L. Katz. 2022. “Predictors of Sleep Disordered Breathing in Children with Down Syndrome: A Systematic Review and Meta-Analysis.” *European Respiratory Review: An Official Journal of the European Respiratory Society*, sleep disordered breathing (SDB), Down syndrome, predictors, obstructive sleep apnea (OSA), systematic review, meta-analysis, neurocognitive morbidity., 31 (164): 220026. <https://doi.org/10.1183/16000617.0026-2022>.
- “Hemiarthroplasty for Hip Fracture in Down Syndrome: A Retrospective Series of Five Cases.” n.d., Down syndrome, hip fracture, hemiarthroplasty, orthopedic surgery, retrospective study., Accessed December 7, 2024. <https://www.hipandpelvis.or.kr/journal/view.html?uid=783&vmd=Full>.
- Howard, Javier J. M., Kathleen M. Sarber, Wenwen Yu, David F. Smith, Raisa O. Tikhtman, Narong Simakajornboon, and Stacey L. Ishman. 2020. “Outcomes in Children with down Syndrome and Mild Obstructive Sleep Apnea Treated Non-Surgically.” *The Laryngoscope*, Obstructive sleep

apnea, mild, observation, infant, pediatric, children, medication., 130 (7): 1828–35. <https://doi.org/10.1002/lary.28325>.

- Kodish, E., and L. Cuttler. 1996. “Ethical Issues in Emerging New Treatments Such as Growth Hormone Therapy for Children with Down Syndrome and Prader-Willi Syndrome.” *Current Opinion in Pediatrics*, Growth hormone therapy, ethical issues, Down syndrome, Prader-Willi syndrome, pediatric treatments, 8 (4): 401–5. <https://doi.org/10.1097/00008480-199608000-00018>.

- Lakshmi, K. T., R. H. Surekha, B. Srikanth, and A. Jyothy. 2008. “University of Coimbra, Faculty of Medicine, Portugal.” *Singapore Medical Journal*, University of Coimbra, Faculty of Medicine, medical research, Portugal, academic institution, clinical studies, medical education., 49 (7): 561–64.

- Lee, P., R. Bhansali, S. Izraeli, N. Hijjiya, and J. D. Crispino. 2016. “The Biology, Pathogenesis and Clinical Aspects of Acute Lymphoblastic Leukemia in Children with Down Syndrome.” *Leukemia*, Down syndrome, acute lymphoblastic leukemia (ALL), CRLF2, JAK2, genetics., 30 (9): 1816–23. <https://doi.org/10.1038/leu.2016.164>.

- Lucas, Patricia, Kristin Liabo, and Helen Roberts. 2002. “Do Behavioural Treatments for Sleep Disorders in Children with Down’s Syndrome Work?” *Archives of Disease in Childhood*, Behavioral treatments, sleep disorders, Down syndrome, sleep training, nighttime behavior., 87 (5): 413–14. <https://doi.org/10.1136/adc.87.5.413>.

- Lukowski, Angela F., Helen M. Milojevich, and Lauren Eales. 2019. “Cognitive Functioning in Children with Down Syndrome: Current Knowledge and Future Directions.” *Advances in Child Development and Behavior*, Down syndrome, cognitive development, intellectual disability, memory, future research., 56:257–89. <https://doi.org/10.1016/bs.acdb.2019.01.002>.

- Marqui, Alessandra Bernadete Trovó de, and Maria de Fátima Borges. 2024. “(In)Fertility in the Down Syndrome.” *Revista Da Associação Médica Brasileira*, Down syndrome, infertility, reproduction, pregnancy outcomes, fertility counseling, 70 (August):e20240537. <https://doi.org/10.1590/1806-9282.20240537>.

- Næss, Kari-Anne B., Monica Melby-Lervåg, Charles Hulme, and Solveig-Alma Halaas Lyster. 2012. “Reading Skills in Children with Down Syndrome: A Meta-Analytic Review.” *Research in Developmental Disabilities*, Down syndrome, reading skills, meta-analysis, decoding, language comprehension, phonological awareness, 33 (2): 737–47. <https://doi.org/norway>.

- Nicek, Anna, Nasreen Talib, Daniel Lovell, Chelsey Smith, Mara L. Becker, and Jordan T. Jones. 2020. “Assessment and Treatment of Down Syndrome-Associated Arthritis: A Survey of Pediatric Rheumatologists.” *Pediatric Rheumatology Online Journal*, Down syndrome, arthritis, pediatric rheumatologists, juvenile idiopathic arthritis, treatment, diagnosis, 18 (1): 57. <https://doi.org/10.1186/s12969-020-00445-6>.

- Nordstrøm, Marianne, Kjetil Retterstøl, Sigrun Hope, and Svein Olav Kolset. 2020. “Nutritional Challenges in Children and Adolescents with Down Syndrome.” *The Lancet. Child & Adolescent Health*, Down syndrome, childhood obesity, Mediterranean diet, dyslipidemia, physical activity, KIDMED questionnaire, 4 (6): 455–64. [https://doi.org/10.1016/S2352-4642\(19\)30400-6](https://doi.org/10.1016/S2352-4642(19)30400-6).

- Okazaki, Fumiko, Hiroyuki Wakiguchi, Yuno Korenaga, Kazumasa Takahashi, Hiroki Yasudo, Ken Fukuda, Mototsugu Shimokawa, and Shunji Hasegawa. 2022. “Food Protein-Induced Enterocolitis Syndrome in Children with Down Syndrome: A Pilot Case-Control Study.” *Nutrients*, FPIES, Down syndrome, food allergy, gastrointestinal disorder, case-control study., 14 (2): 388. <https://doi.org/10.3390/nu14020388>.

- Poudel, Aashis. 2023. “Insights into Children with Down Syndrome: A Medical Student’s Perspective.” *JNMA; Journal of the Nepal Medical Association*, Developmental disabilities, Down syndrome, holistic health, medical student., 61 (264): 680–82. <https://doi.org/10.31729/jnma.8244>.

- Prena, Kelsey, and John L. Sherry. 2018. “Parental Perspectives on Video Game Genre Preferences and Motivations of Children with Down Syndrome.” *Journal of Enabling Technologies*, Down syndrome, children, video games, parental perspectives, genre preferences, motivations, , March. <https://www.emerald.com/insight/content/doi/10.1108/jet-08-2017-0034/full/html>.

- Ray, Anirban, Tiffany Rene Oliver, Pinku Halder, Upamanyu Pal, Sumantra Sarkar, Supratim Dutta, and Sujay Ghosh. 2018. “Risk of Down Syndrome Birth: Consanguineous Marriage Is Associated with Maternal meiosis-II Nondisjunction at Younger Age and without Any Detectable Recombination Error.” *American Journal of Medical Genetics Part A*, Down syndrome,

consanguineous marriage, meiosis-II nondisjunction, maternal age, recombination errors, trisomy 21., 176 (11): 2342–49. <https://doi.org/10.1002/ajmg.a.40511>.

- Rosso, Mattia, Ellen Fremion, Stephanie L. Santoro, Nicolas M. Oreskovic, Tanuja Chitnis, Brian G. Skotko, and Jonathan D. Santoro. 2020. “Down Syndrome Disintegrative Disorder: A Clinical Regression Syndrome of Increasing Importance.” *Pediatrics*, Down syndrome, disintegrative disorder, regression, catatonia, psychiatric conditions, developmental delay, adolescence, 145 (6): e20192939. <https://doi.org/10.1542/peds.2019-2939>.

- Sheets, Kathryn B., Robert G. Best, Campbell K. Brasington, and Madeleine C. Will. 2011. “Balanced Information about Down Syndrome: What Is Essential?” *American Journal of Medical Genetics Part A*, Down syndrome, diagnosis, information delivery, parental support, healthcare professionals., 155 (6): 1246–57. <https://doi.org/10.1002/ajmg.a.34018>.

- Siriwardhana, Leon S., Gillian M. Nixon, Margot J. Davey, Dwayne L. Mann, Shane A. Landry, Bradley A. Edwards, and Rosemary S. C. Horne. 2021. “Children with down Syndrome and Sleep Disordered Breathing Display Impairments in Ventilatory Control.” *Sleep Medicine*, Down syndrome, sleep disordered breathing, ventilatory control, loop gain, polysomnography., 77 (January):161–69. <https://doi.org/10.1016/j.sleep.2020.12.005>.

- Skotko, B. G. 2009. “With New Prenatal Testing, Will Babies with Down Syndrome Slowly Disappear?” *Archives of Disease in Childhood*, Prenatal testing, Down syndrome, abortion, ethical concerns, prenatal diagnosis, 94 (11): 823–26. <https://doi.org/10.1136/adc.2009.166017>.

- Souza, Denise E. De, and Athena Vongalis-Macrow. 2024. “I’m Trying to Mix, but It’s Really Hard to Talk and Explain Ideas: Inclusion of Students With Down Syndrome in Higher Education.” *Including Voices*, Inclusion, higher education, Down syndrome, accessibility, barriers, support systems, 23 (June):177–89.

- Souza Oliveira, Renato, João Quadrado Gil, Andreia Rosa, Maria João Quadrado, and Mauro Campos. 2024. “Scheimpflug Tomographic Indices for Classifying Normal, Down Syndrome and Clinical Keratoconus in Pediatric Patients.” *Diagnostics*, keratoconus, Down syndrome, corneal tomography, pediatric, Pentacam, diagnostic indices, 14 (17): 1932. <https://doi.org/10.3390/diagnostics14171932>.

- Stefferud, Marte Johanne, Anne Grethe Einang, and Claus Klingenberg. 2021. “Parents of Children with Down Syndrome and Their Experiences with the Healthcare Services.” *Tidsskrift for Den Norske Laegeforening: Tidsskrift for Praktisk Medicin, Ny Raekke*, Down syndrome, healthcare services, parental experiences, support, early years, healthcare needs., 141 (September). <https://doi.org/10.4045/tidsskr.21.0024>.

- Tenenbaum, Ariel, Rana N. Hanna, Diana Averbuch, Isaiah D. Wexler, Maor Chavkin, and Joav Merrick. 2014. “Hospitalization of Children with down Syndrome.” *Frontiers in Public Health*, Down syndrome, children, hospitalization, respiratory diseases, ICU care, comorbidities, mortality rates., 2:22. <https://doi.org/10.3389/fpubh.2014.00022>.

- Thorneycroft, Ryan. 2022. “Prenatal Testing, Down Syndrome, and Selective Termination: A (Critical) Criminology of Genocide?” *Diversity in Criminology and Criminal Justice Studies*, Prenatal testing, Down syndrome, selective termination, genocide, disability ethics, reproductive choices., 27 (May):167–81.

- Van Riper, M., and W. I. Cohen. 2001. “Caring for Children with Down Syndrome and Their Families.” *Journal of Pediatric Health Care: Official Publication of National Association of Pediatric Nurse Associates & Practitioners*, Down syndrome, pediatric care, family support, early intervention., 15 (3): 123–31. <https://doi.org/10.1067/mpn.2001.110627>.

- Walton, Catherine, and Mike Kerr. 2015. “Down Syndrome: Systematic Review of the Prevalence and Nature of Presentation of Unipolar Depression.” *Advances in Mental Health and Intellectual Disabilities*, Down syndrome, unipolar depression, psychiatric disorders, prevalence, intellectual disability, mood disorders, 9 (4): 151–62. <https://doi.org/10.1108/AMHID-11-2014-0037>.

- Wang, Xin, Qian Yang, Xueyan Zhou, C. Dirk Keene, Alexey G. Ryazanov, and Tao Ma. 2024. “Suppression of eEF2 Phosphorylation Alleviates Synaptic Failure and Cognitive Deficits in Mouse Models of Down Syndrome.” *Alzheimer’s & Dementia*, Down syndrome, cognitive deficits, synaptic failure, eEF2 phosphorylation, eEF2K inhibition, mouse models., 20 (8): 5357–74. <https://doi.org/USA>.

- Weger, Christine de, Nienke Boonstra, and Jeroen Goossens. 2019. “Effects of Bifocals on Visual Acuity in Children with Down Syndrome: A Randomized Controlled Trial.” *Acta*

Ophthalmologica, Down syndrome, bifocals, visual acuity, randomized controlled trial, 97 (4): 378–93. <https://doi.org/10.1111/aos.13944>.

- Wohlfert, Abigail J., Jeremiah Phares, and Ann-Charlotte Granholm. 2024. “The mTOR Pathway: A Common Link Between Alzheimer’s Disease and Down Syndrome.” *Journal of Clinical Medicine*, mTOR pathway, Alzheimer’s disease, Down syndrome, PI3K/Akt signaling, autophagy, tau phosphorylation, amyloid-beta, 13 (20): 6183. <https://doi.org/10.3390/jcm13206183>.

- Zemel, Babette S., Mary Pipan, Virginia A. Stallings, Waynitra Hall, Kim Schadt, David S. Freedman, and Phoebe Thorpe. 2015. “Growth Charts for Children With Down Syndrome in the United States.” *Pediatrics*, Down syndrome, growth charts, children, weight-for-length, BMI, pediatrics, growth assessment, developmental milestones., 136 (5): e1204-1211.

