# Music therapy for children going through haematopoietic stem cell transplantation

Lena Uggla & Lars Ole Bonde

#### Introduction

Paediatric haematopoietic stem cell transplantation (HSCT) is a well-known treatment for aggressive leukemias, advanced haematopoietic and metabolic diseases (Miano et al., 2007). Medical developments in HSCT procedures have led to increased use of HSCT, and overall survival has improved in the paediatric population (Remberger et al., 2011). During the initial three to six months the HSCT procedure is very intense, including a period of isolation for 4–6 weeks, mainly due to the risk of infection. The child is monitored frequently after the initial hospitalisation in order to early detect infections etc. but also graft-versus-host disease (GVHD) or relapse of the disease.

The treatment affects the whole body and has a major impact on the child's and the parents' psychological wellbeing as well as their health-related quality of life (HRQoL) (Packman, Weber, Wallace & Bugescu, 2010). The lowest ratings of the child's HRQoL are noted one and three months after the HSCT (Rodgers, Wills-Bagnato, Sloane & Hockenberry, 2015). It takes approximately one to three years to return to the same level of HRQoL as before HSCT (Tanzi, 2011; Tremolada, Bonichini, Pillon, Messina & Carli, 2009).

Going through HSCT can be life-threatening. In terms of psychological reactions, posttraumatic stress disorder (PTSD) and traumatic stress symptoms as well as neurocognitive dysfunctions have been reported in HSCT survivors (Buchbinder et al., 2018; Stuber, Nader, Yasuda, Pynoos & Cohen, 1991). Earlier research has also reported high levels of stress and depressive symptoms in families of children going through HSCT (Phipps, Dunavant, Lensing & Rai, 2005).

#### Why music and music therapy?

Music affects the whole brain and the neurochemical systems of reward, motivation and pleasure, and can reduce stress levels and strengthen social attachments (Chanda & Levitin, 2013). Music evokes and affects emotions. The use of music in emotional regulation is supported by behavioural and neural evidence due to music's function in early infant-parent bonding and its developmental fitness (Sena Moore & Hanson-Abromeit, 2015). Musical activations also have an effect on different biomarkers, e.g. the stress hormone cortisol and reducing increases in blood glucose. Familiar music, singing, creating and improvising are other factors that seem to have an impact on emotional regulation and reduce activity in the amygdala (Fancourt, Ockelford & Belai, 2014; Finn & Fancourt, 2018; Moore, 2013).

Music therapy is used in paediatric healthcare with the goal of helping children through serious experiences and supporting health (Bradt, 2012). Music therapy is a relational therapy, and one main objective is to increase the patient's experiences and intersubjective knowledge by being involved and relating through music (Trondalen, 2016).

Previous research reported increased well-being and decreased procedural pain after music interventions for children with cancer (Barrera, Rykov & Doyle, 2002; Nguyen, Nilsson, Hellstrom & Bengtson, 2010). Music therapy in the HSCT context for children showed reduced levels of anxiety and, in the young adult population, improved coping and social integration were reported (Robb et al., 2014; Robb & Ebberts, 2003).

# Study design and music therapy protocol

From February 2013 to November 2017 a randomised study was conducted at the Karolinska University Hospital, Sweden. Thirty-eight patients from 2 months to 17 years of age were included in the study. Music therapy was performed twice a week for 4–6 weeks in the music therapy group. The patients were hospitalised until donor engraftment. After engraftment the children were monitored in an outpatient paediatric ward at the hospital. The children in the control group were offered music therapy after discharge in the outpatient ward, twice a week for 4–6 weeks.

The music therapy intervention included both expressive and receptive methods (MacDonald, Kreutz & Mitchell, 2012). The session took place in the child's hospital room, and the child was invited to play different musical instruments, sing and listen to music with the music therapist.

The parents and siblings could also participate. The music therapy intervention was designed to build a trusting relationship between the child and the music therapist. The session had the goal of being flexible and varied in order to provide a holding structure to benefit both children and parents so that they could stay emotionally regulated (Fosha, Siegel & Solomon, 2009).

The aims of our study were:

- To evaluate physiological endpoints such as heart rate, blood pressure and saturation as well as evaluation of pain and mood between treatment groups during HSCT. The music therapy group received music therapy twice a week, and the control group received supportive conventional treatment.
- To compare HRQoL questionnaires at admission, discharge and after 6 months between music therapy group and control group.
- To explore the subjective experiences and memories of interactions between children, parents and music therapist during music therapy interventions.

### Results

We have published three articles based on our research project. In our first article, which included 21 patients, 0-16 years of age, we reported on the children's heart rates, blood pressure and saturation in connection with music therapy (Uggla et al., 2016). The children's physiological parameters in the morning were noted in both the music therapy group and the control group and were then compared with corresponding parameters that were recorded in the evening. The children's heart rates in the music therapy group were significantly lower (p < 0.001) in the evening 4–8 hours after the music therapy intervention in comparison to the control group.

In our second article, which included 29 patients, 0-17 years of age, we reported that the children's self-reported health-related quality of life (HRQoL) improved after the music therapy intervention. In the music therapy group, the physical functioning improved at the time of discharge (adjusted p = 0.04), and the control group showed improved results after the music therapy interventions in all domains. (p= 0.015) (Uggla, Bonde, Hammar, Wrangsjo & Gustafsson, 2018).

Our third article was a qualitative study using the collaborative interview format as a data collection method, examining the families' experiences and memories of the interaction in the music therapy session. One of the questions was: what was it like to experience music therapy? Six families were included, and child, parents and music therapist were interviewed at

the same time by an independent psychologist who also performed the analysis. The analysis emerged in three themes: experiences of competence and recognition of the self; experiences of interactive affect regulation as potential for change; experiences of the importance of the therapeutic relationship (Uggla, Mårtenson Blom, Bonde, Gustafsson & Wrangsjö, 2019).

#### Discussion

The aim of the doctoral study was to evaluate and explore music therapy in the particular HSCT context. Heightened heart rate was reported to be related to increased levels of distress, and elevated heart rate after an accident may predict symptoms of PTSD 6 months later for children and adolescents (De Young, Kenardy & Spence, 2007; Morris, Hellman, Abelson & Rao, 2016). Previous research has reported symptoms of stress in children, parents and siblings after HSCT. The significantly decreased heart rate levels in the evening in the music therapy group may indicate decreased levels of stress due to decreased activation in the amygdala through emotional regulation.

The increasing physical functioning reported by the children at discharge in the music therapy group and the overall increased HRQoL in the control group after music therapy at 6 months follow-up indicate the importance of the music therapy intervention. The musical elements – melody, rhythm, movement and dynamic shifts in intensity – have the potential to facilitate an intersubjective regulation experience. Shared affects and intentions in music therapy are cross-modal and do not need to be explained in words. Affect attunement through musical experiences in music therapy gives the participants an opportunity to interact and thereby influence the interplay. This could be seen as important to these children and parents, whose current life circumstances greatly reduce their opportunity to influence their situation and the challenging medical treatment.

# Conclusions

Music therapy developed into a significant and helpful experience for the participants, an important factor in coping and managing the treatment period at the hospital. The combination of reduced heart rate values four to eight hours after the intervention in the music therapy group and the improved HRQoL reported by both groups suggests that music therapy can be an effective, complementary intervention during and after HSCT.

#### References

- Barrera, M. E., Rykov, M. H. & Doyle, S. L. (2002). The effects of interactive music therapy on hospitalized children with cancer: A pilot study. *Psychooncology*, 11(5), 379–388. https://doi.org/10.1002/pon.589
- Bradt, J. (Ed.) (2012). *Guidelines for music therapy practice in pediatric care*. Dallas, TX: Barcelona.
- Buchbinder, D., Kelly, D. L., Duarte, R. F., Auletta, J. J., Bhatt, N., Byrne, M., . . . Shaw, B.
  E. (2018). Neurocognitive dysfunction in hematopoietic cell transplant recipients:
  Expert review from the late effects and Quality of Life Working Committee of the CIBMTR and complications and Quality of Life Working Party of the EBMT. *Bone Marrow Transplant*, *53*(5), 535–555. https://doi.org/10.1038/s41409-017-0055-7
- Chanda, M. L. & Levitin, D. J. (2013). The neurochemistry of music. *Trends in Cognitive Sciences*, *17*(4), 179–193. https://doi.org/10.1016/j.tics.2013.02.007
- De Young, A. C., Kenardy, J. A. & Spence, S. H. (2007). Elevated heart rate as a predictor of PTSD six months following accidental pediatric injury. *J Trauma Stress*, 20(5), 751–756. https://doi.org/10.1002/jts.20235
- Fancourt, D., Ockelford, A. & Belai, A. (2014). The psychoneuroimmunological effects of music: A systematic review and a new model. *Brain Behav Immun*, 36, 15–26. https://doi.org/10.1016/j.bbi.2013.10.014
- Finn, S. & Fancourt, D. (2018). The biological impact of listening to music in clinical and nonclinical settings: A systematic review. *Prog Brain Res*, 237, 173–200. https://doi. org/10.1016/bs.pbr.2018.03.007
- Fosha, D., Siegel, D. J. & Solomon, M. F. (2009). *The healing power of emotion : Affective neuroscience, development, and clinical practice.* New York: Norton.
- MacDonald, R. A. R., Kreutz, G. & Mitchell, L. (2012). *Music, health, and wellbeing*. Oxford: Oxford University Press.
- Miano, M., Labopin, M., Hartmann, O., Angelucci, E., Cornish, J., Gluckman, E., ...
  Marrow, T. (2007). Haematopoietic stem cell transplantation trends in children over the last three decades: a survey by the paediatric diseases working party of the European Group for Blood and Marrow Transplantation. *Bone Marrow Transplant, 39*(2), 89–99. https://doi.org/10.1038/sj.bmt.1705550
- Moore, K. S. (2013). A systematic review on the neural effects of music on emotion regulation: Implications for music therapy practice. *J Music Ther*, *50*(3), 198–242. Retrieved from http://www.ncbi.nlm.nih.gov/pubmed/24568004
- Morris, M. C., Hellman, N., Abelson, J. L. & Rao, U. (2016). Cortisol, heart rate, and blood pressure as early markers of PTSD risk: A systematic review and meta-analysis. *Clin Psychol Rev*, 49, 79–91. https://doi.org/10.1016/j.cpr.2016.09.001

- Nguyen, T. N., Nilsson, S., Hellstrom, A. L. & Bengtson, A. (2010). Music therapy to reduce pain and anxiety in children with cancer undergoing lumbar puncture: A randomized clinical trial. *J Pediatr Oncol Nurs*, 27(3), 146–155. https://doi.org/10.1177/1043454209355983
- Packman, W., Weber, S., Wallace, J. & Bugescu, N. (2010). Psychological effects of hematopoietic SCT on pediatric patients, siblings and parents: A review. *Bone Marrow Transplant*, 45(7), 1134–1146. https://doi.org/10.1038/bmt.2010.74
- Phipps, S., Dunavant, M., Lensing, S. & Rai, S. N. (2005). Psychosocial predictors of distress in parents of children undergoing stem cell or bone marrow transplantation. *J Pediatr Psychol*, 30(2), 139–153. https://doi.org/10.1093/jpepsy/jsi002
- Remberger, M., Ackefors, M., Berglund, S., Blennow, O., Dahllof, G., Dlugosz, A., . . . Ringden, O. (2011). Improved survival after allogeneic hematopoietic stem cell transplantation in recent years. A single-center study. *Biol Blood Marrow Transplant*, *17*(11), 1688–1697. https://doi.org/10.1016/j.bbmt.2011.05.001
- Robb, S. L., Burns, D. S., Stegenga, K. A., Haut, P. R., Monahan, P. O., Meza, J., . . . Haase, J. E. (2014). Randomized clinical trial of therapeutic music video intervention for resilience outcomes in adolescents/young adults undergoing hematopoietic stem cell transplant: A report from the Children's Oncology Group. *Cancer*, 120(6), 909–917. https://doi.org/10.1002/cncr.28355
- Robb, S. L. & Ebberts, A. G. (2003). Songwriting and digital video production interventions for pediatric patients undergoing bone marrow transplantation, part II: An analysis of patient-generated songs and patient perceptions regarding intervention efficacy. *Journal of Pediatric Oncology Nursing*, 20(1), 16–25. https://doi.org/10.1053/jpon.2003.4
- Rodgers, C., Wills-Bagnato, P., Sloane, R. & Hockenberry, M. (2015). Health-related quality of life among children and adolescents during hematopoietic stem cell transplant recovery. *Journal of Pediatric Oncology Nursing*, 32(5), 329–336. https://doi.org/10.1177/1043454214563413
- Sena Moore, K. & Hanson-Abromeit, D. (2015). Theory-guided therapeutic function of music to facilitate emotion regulation development in preschool-aged children. *Frontiers in Human Neuroscience*, 9. https://doi.org/10.3389/fnhum.2015.00572
- Stuber, M. L., Nader, K., Yasuda, P., Pynoos, R. S. & Cohen, S. (1991). Stress responses after pediatric bone marrow transplantation: Preliminary results of a prospective longitudinal study. J Am Acad Child Adolesc Psychiatry, 30(6), 952–957. https://doi.org/10.1097/00004583-199111000-00013
- Tanzi, E. M. (2011). Health-related quality of life of hematopoietic stem cell transplant childhood survivors: State of the science. *J Pediatr Oncol Nurs*, 28(4), 191–202. https://doi.org/10.1177/1043454211408100

Tremolada, M., Bonichini, S., Pillon, M., Messina, C., & Carli, M. (2009). Quality of life and psychosocial sequelae in children undergoing hematopoietic stem-cell transplantation: A review. *Pediatr Transplant*, *13*(8), 955–970. https://doi.org/10.1111/j.1399-3046.2009.01203.x

Trondalen, G. (2016). *Relational music therapy : an intersubjective perspective*. Dallas, TX: Barcelona.

- Uggla, L., Bonde, L. O., Hammar, U., Wrangsjo, B. & Gustafsson, B. (2018). Music therapy supported the health-related quality of life for children undergoing haematopoietic stem cell transplants. *Acta Paediatr*, *107*(11), 1986–1994. https://doi.org/10.1111/apa.14515
- Uggla, L., Bonde, L. O., Svahn, B. M., Remberger, M., Wrangsjo, B. & Gustafsson, B. (2016). Music therapy can lower the heart rates of severely sick children. *Acta Paediatr*, 105(10), 1225–1230. https://doi.org/10.1111/apa.13452
- Uggla, L., Mårtenson Blom, K., Bonde, L. O., Gustafsson, B. & Wrangsjö, B. (2019). An explorative study of qualities in interactive processes with children and their parents in music therapy during and after pediatric hematopoietic stem cell transplantation. *Medecines*, 6(1), 28. Retrieved from http://www.mdpi.com/ 2305-6320/6/1/28

Lena Uggla<sup>a\*</sup> & Lars Ole Bonde<sup>b</sup> <sup>a</sup>Karolinska Institutet, Stockholm, Sweden, <sup>b</sup>independent senior researcher (former Aalborg University, Denmark)

\*Corresponding author Lena Uggla *Karolinska Institutet* Stockholm Sweden tel. +46705298010 e-mail: lena.uggla@ki.se