

# MANUELA SPADEA, MD



## PRESENT POSITION

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**University of Turin, Regina Margherita Children's Hospital, Turin (Italy)** *07/2021-ongoing*

**Division:** Department of Pediatric and Public Health Sciences- Pediatric Onco-hematology, Stem Cell Transplant Service and Cell Therapy Division

**Position:** Assistant Professor of Pediatrics

**Co-investigator** in several phase I-II-III Clinical Trials *07/2019 - ongoing*

**Co-PI** of NCT05523336 Trial *08/2022- ongoing*

**Study Coordinator** in Turin Metropolitan Transplant Center *05/2021- ongoing*

## PREVIOUS PROFESSIONAL POSITIONS

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**Regina Margherita Children's Hospital, Turin (Italy)** *12/2021- 06/2022*

**Division:** Pediatric Onco-hematology, Stem Cell Transplant Service and Cell Therapy

**Position:** Pediatric Oncologist Consultant

## CLINICAL AND RESEARCH EXPERIENCE ABROAD

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**Boston Children's Hospital/Dana Farber Cancer Institute, Harvard Medical School, Boston (MA,USA)** *03/2019- 04/2019*

**Division:** Stem Cell Transplant Service

**Position:** Clinical observer

**Tutor:** Prof. Leslie Lehmann, MD.

**Boston Children's Hospital/Dana Farber Cancer Institute, Harvard Medical School, Boston (MA,USA)** *04/2019 - 06/2019*

**Gene Therapy Program, Dana Farber/ Boston Children's Cancer and Blood Disorders Center**

**Position:** Visiting Scientist

**Research Project:** Development of novel treatments for severe diseases using genetic correction approaches such as viral vectors and CRISPR-Cas9 based gene editing

**Tutor:** Prof. Christian Brendel, PhD.

## OTHER RESEARCH EXPERIENCES

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**Centro di formazione e simulazione neonatale NINA- (Pisa, Italy)** *06/2020-10/2020*

**Position:** Research Assistant

**Main projects:** Simulation-based medical education in pediatrics; artificial intelligence applied to non-technical skills in resuscitation teams

**Tutor:** Armando Cuttano, MD

## MEDICAL LICENSURE

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National Professional Qualification as Physician - Albo TO 23859

02/2016

## EDUCATION

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**Regina Margherita Children's Hospital, University of Turin (Italy)** 11/2016 - 11/2021

**Residency in Pediatrics**

**Final Mark:** 70/70 cum laude

**Thesis Title:** *Multivariate Analysis of Immune Reconstitution and Relapse Risk Scoring in Children Receiving Allogeneic Stem Cell Transplantation for Acute Leukemias*

**Tutors:** *Ugo Ramenghi, Prof.; Franca Fagioli, Prof.*

**University of Pisa, Pisa (Italy)**

09/2009 - 10/2015

**Master's Degree in Medicine and Surgery**

**Final Mark:** 110/110 cum laude

**Thesis Title:** *Il trapianto di cellule staminali emopoietiche da donatore familiare aploidentico: analisi delle strategie innovative e della ricostituzione immunologica in pazienti dell'U.O. di Oncoematologia Pediatrica di Pisa*

**Thesis Title (English):** *Haploidentical stem cell transplantation: an analysis of innovative strategies and immune reconstitution in patients of the pediatric hemato-oncology department of Pisa University Hospital.*

**Tutors:** *Maria Cristina Menconi, MD; Claudio Favre, MD*

**Scientific High School E. Majorana, Girifalco (CZ), Italy**

09/2004 - 06/2009

**Scientific High School Diploma**

**Final Mark:** 100/100 cum laude

**Music Conservatory F.Torrefranca, Vibo Valentia (VV), Italy**

2007/2008

**Solfeggio and lower level lyric vocal Diploma**

**Final Mark:** 9/10

## PUBLICATIONS

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[1] Saglio F, Berger M, **Spadea M**, Pessolano R, Carraro F, Barone M, and Fagioli F(2020). *Haploidentical HSCT with post transplantation cyclophosphamide versus unrelated donor HSCT in pediatric patients affected by acute leukemia*. Bone Marrow Transplant. 2021 Mar;56(3):586-595. doi: 10.1038/s41409-020-01063-2. Epub 2020 Sep 23. PMID: 32968215.

[2] **Spadea M**, Carraro F., Saglio F., Pessolano R., Berger M., Vassallo E. and Fagioli F. *Successfully treated severe COVID-19 and invasive aspergillosis in early hematopoietic cell transplantation setting*. Transplant Infectious Disease (2020): e13470.

[3] Del Borrello G, Giraudo I, Bondone C, Denina M, Garazzino S, Linari C, Mignone F, Pruccoli G, Scolfaro C, **Spadea M**, Pollio B, Saracco P. *SARS-CoV-2 associated coagulopathy and thromboembolism prophylaxis in hospitalised children: a single center observational study*, J Thromb Haemost. 2021 Feb;19(2):522-530. doi: 10.1111/jth.15216. Epub 2020 Dec 23. PMID: 33305475.

[4] **Spadea M**, Ciantelli M, Fossati N, Cuttano A. *Enhancing the future of simulation-based education in pediatrics*. Ital J Pediatr. 2021 Feb 17;47(1):36. doi: 10.1186/s13052-021-00989-7. PMID: 33596954; PMCID: PMC7888690.

- [5] Quarello P, Perotti D, Carli D, Giorgio E, Sirchia F, Brusco A, Ferrero GB, Mussa A, **Spadea M**, Ciceri S, Spreafico F, Fagioli F. *Wilms tumour occurring in a patient with osteopathia striata with cranial sclerosis: A still unsolved biological question.* [published online ahead of print, 2021 May 24]. *Pediatr Blood Cancer*. 2021;e29132. doi:10.1002/pbc.29132
- [6] Zucchetti G, Ciappina S, Bellini S, Dionisi Vici M, **Spadea M**, Biasin E, Fagioli F. *The Creation of a Transition Protocol Survey for Childhood, Adolescent, and Young Adult Cancer Survivors in Transition from Pediatric to Adult Health Care in Italy.* *J Adolesc Young Adult Oncol*. 2022 Jan 24. doi: 10.1089/jayao.2021.0037. Epub ahead of print. PMID: 35076255.
- [7] Ceolin V, Cenna R, Resente F, **Spadea M**, Fagioli F, Bertorello N. *Guillain-Barre' syndrome after bortezomib therapy in a child with relapsed acute lymphoblastic leukemia,* [published online ahead of print, 2021 Aug 9]. *Pediatr Hematol Oncol*. 2021;1-6. doi:10.1080/08880018.2021.1959691
- [8] **Spadea M**, Saglio F, Tripodi SI, Menconi M, Zecca M, Fagioli F. *Multivariate analysis of immune reconstitution and relapse risk scoring in children receiving allogeneic stem cell transplantation for acute leukemias.* *Transplant Direct*. 2021;7(11):e774. Published 2021 Oct 6. doi:10.1097/TXD.0000000000001226
- [9] **Spadea M**, Saglio F, Pessolano R, Opramolla A, Calvo PL, Fagioli F. *Diagnosing acute intestinal graft-versus-host disease by a non-invasive method: transabdominal ultrasonography and colour doppler imaging.* *Lancet*. 2021;398(10306):1170. doi:10.1016/S0140-6736(21)01918-8
- [10] **Spadea M**, Quarello P, Saglio F, Pedace L, Fagioli F. *GATA2-related myeloid neoplasms in pediatrics: where do we stand?* [published online ahead of print, 2021 Oct 20]. *Pediatr Hematol Oncol*. 2021;1-5. doi:10.1080/08880018.2021.1987600
- [11] Berger M, Barone M, **Spadea M.**, Saglio F, Pessolano R, Fagioli F. *HSCT with mismatched unrelated donors: Bone marrow versus peripheral blood stem cells sources in pediatric patients.* *Pediatr Transplant*. 2022 Jan 28:e14233. doi: 10.1111/petr.14233. Epub ahead of print. PMID: 35092128.
- [12] Berger M, Barone M, Carnevale-Schianca F, De Gobbi M, Nicoli P, Caravelli D, Cilloni D, Paruzzo L, **Spadea M.**, Mareschi K, et al. *HSCT with Mismatched Unrelated Donors (MMUD): A Comparison of Different Platforms for GvHD Prophylaxis.* *Transplantation* 2022, 3, 51–67. <https://doi.org/10.3390/transplantation3010006>
- [13] Mareschi K, Ferrero I, Pinnetta G, Leone M, Marini E, Adamini A, Banche Niclot AGS, **Spadea M**, Fagioli F. *Terapie cellulari. Le potenzialità terapeutiche delle cellule staminali mesenchimali.* *Giornale dell'accademia di Medicina di Torino, Anno CLXXXIII, 2020.*
- [14] Mareschi K, Marini E, Banche Niclot AGS, Barone M, Pinnetta G, Adamini A, **Spadea M**, Labanca L, Lucania G, Ferrero I, Fagioli F. *A new human platelet lysate for mesenchymal stem cell production compliant with good manufacturing practice conditions.* *Int. J. Mol. Sci.*, March 2022
- [15] **Spadea M**, Saglio F, Opramolla A, Rigazio C, Cisarò F, Berger M, Quarello P, Calvo PL, Fagioli F. *Non-Invasive Diagnosis of Pediatric Intestinal Graft-Versus-Host Disease: A Case Series.* *Transplantation*. 2022; 3(2):115-123. <https://doi.org/10.3390/transplantation3020012>
- [16] Quarello, P.; Carli, D.; BIASONI, D.; Gerocarni Nappo, S.; Morosi, C.; Cotti, R.; Garelli, E.; Zucchetti, G.; **Spadea, M.**; Tirtei, E.; et al. *Implications of an Underlying Beckwith–Wiedemann Syndrome for Wilms Tumor Treatment Strategies.* *Cancers* 2023, 15, 1292.
- [17] **Spadea, M.**, Saglio F, Ceolin V, Barone M, Quarello P, Fagioli F. *Immune-mediated cytopenias (IMCs) after HSCT for pediatric non-malignant disorders: epidemiology, risk factors, pathogenesis, treatment.* *European Journal Of Pediatrics*, 2023, accepted, in press

- [1] Menconi, M., **Spadea, M.**, Giannelli, R., Bernasconi, S., Bulleri, M., Casazza, G., ... ,and Favre, C. (2016). *Haploidentical Transplantation (haplo Hsct): high survival, low incidence of graft-versus-host-disease and fast immune reconstitution using Cd3 or Tcr $\alpha\beta$  and Cd19 depletion in the experience of Pediatric Hematology Oncology Department of Pisa*. Bone Marrow Transplantation, 51, S575-S574.
- [2] **Spadea M.**, Vallerio S., Basso E, Bertin D., Morra I., Peretta P., Fagioli F. (2018). *Sindrome da predisposizione a tumori rabdoidi (RTPS) in paziente con neoplasie rabdoidi metacrone (Tumore rabdoide renale, AT/RT)*. Hematology Reports 2018, XLIII AIEOP Congress, Bologna, Italy.
- [3] **Spadea M.**, Saglio F., Berger M., Carraro F., Pessolano R., Ricotti E., Vassallo E., Fagioli F. (2020). *A new multivariate model to describe impact of immune reconstitution on the development of relapse in children undergoing allo-HSCT for hematologic malignancies at day +100*. BONE MARROW TRANSPLANTATION. Vol. 55. No. SUPPL 1. CAMPUS, 4 CRINAN ST, LONDON, N1 9XW, ENGLAND: SPRINGER NATURE, 2020.
- [4] Berger M., Barone M., Saglio F., Carnevale-Schianca F., De Gobbi M., Paruzzo L., Nicoli P., **Spadea M.**, Caravelli D., Vassallo E., Aglietta M., Fagioli F. (2020). *Acute GVHD incidence in mismatched unrelated donor HSCT in paediatric and adult patients: a study from the Turin Metropolitan Transplant Center*. BONE MARROW TRANSPLANTATION. Vol. 55. No. SUPPL 1. CAMPUS, 4 CRINAN ST, LONDON, N1 9XW, ENGLAND: SPRINGER NATURE, 2020.
- [5] **Spadea M.**, Saglio F., Berger M., Carraro F., Pessolano R., Vassallo E., Fagioli F. (2020). *Identificazione precoce di alto rischio di recidiva post-TCSE in relazione alla ricostituzione immunologica mediante analisi delle componenti principali (PCA): una piattaforma per le terapie cellulari profilattiche*. Hematology Reports 2018, XLV AIEOP Congress

## SELECTED PUBLIC SPEECHES AND POSTER PRESENTATIONS

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- [1] **Spadea M.**, Siri B., Basso E (2018). *Rhabdoid tumor predisposition syndrome: a case report*. II Incontro - Gruppo di studio di Neuro-Oncologia e Neurologia dei Trapianti, Pavia, Italy.
- [2] **Spadea M.**, Saglio F., Vassallo E., Ricci F., Fagioli F. (2019). *Un unusual complication in the early post-transplant setting: immune polyneuropathy*. 4th EBMT International Transplant Course, Barcelona, Spain.
- [3] **Spadea M.** F. Saglio, V. Ceolin, G. Del Borrello and Franca Fagioli. The endothelium involvement during CRS: first reported case of CAR-VOD. 5th EBMT-EHA CAR-T cell meeting, Rotterdam, The Netherlands

## COURSES AND CONFERENCES

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**Università degli Studi di Milano** 11/2021-01/2022

**Course:** *Post-Graduate Course in Clinical Research- Corso di Perfezionamento in Ricerca clinica: come disegnare studi randomizzati e osservazionali. Metodi per la valutazione dell'efficacia e sicurezza degli interventi sanitari*

**Main Field:** *Clinical Research, Randomized Controlled Trials, Observational Studies*

**Società italiana Medicina di Emergenza Urgenza pediatrica (SIMEUP)** 06/2021

**Course:** *Pediatric Advanced Life Support-PALS*

**Main Field:** *Pediatrics, Resuscitation*

**Tutor:** *A. Urbino, MD*

I attended several national and international conferences in Pediatrics and Pediatric Oncology and related fields.

## AWARDS

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## OTHER EXPERIENCES

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**Private Practice:** Substitution of Pediatricians in private practice 01/2018 - 11/2021

**Private Practice:** Medical Doctor at Dynamo Camp Onlus, a recreational therapy camp. 12/2018

## SOCIETY MEMBERSHIP

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**EHA:** European Hematology Association; **GITMO:** Gruppo Italiano per il Trapianto di Midollo Osseo, cellule staminali emopoietiche e terapie cellulari; **AIEOP:** Associazione Italiana Ematologia Oncologia Pediatrica; **EBMT:** European Society for Blood and Marrow Transplantation

**Trainee Committee EBMT:** Active Member.

**Infectious Disease and Supportive Therapy Working Group of AIEOP:** Board Member.

## LANGUAGES

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- **Italian**, Native Speaker
- **English**, fluent
- **French**, fluent

## MAIN SKILLS

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**Organization skills:** Research projects scientific management. Group work. Adaptability in multi-cultural working environments.

**Technical skills:** Windows e OS X. Text editor tools LATEX, BIBTEX, Microsoft Office, Open Office; Internet browsing Safari, Internet Explorer, Chrome, Opera; Search engines, PubMed and Google Scholar; familiarity with usage of statistical software R; Good usage skills of SonoSim for FAST (Focused Assessment with Sonography for Trauma).

**Laboratory skills:** restriction digest, bacterial transformation, preparation of plasmid DNA, sanger sequencing, transfection of cell lines for vector production, titration of the vector supernatants on indicator cell lines, analyses of transduced cells by flow cytometry; design and cloning of gRNAs for CRISPR-Cas9 based tagging gene method.

**Other skills:** Lyrical vocalist, solo soprano in concert with choir and orchestra.

*In compliance with the Italian legislative Decree no. 196 dated 30/06/2003, I hereby authorize the recipient of this document to use and process my personal details.*