

## Intravenous pentamidine for *Pneumocystis pneumonia* prophylaxis in children undergoing autologous hematopoietic stem cell transplant

We are writing you with regards to the brief report, "The use of intravenous pentamidine for the prophylaxis of *Pneumocystis pneumonia* in pediatric patients," published in January 2017.<sup>1</sup>

We would like to share our positive experience with intravenous pentamidine (IVP) in prophylaxis of *Pneumocystis jirovecii* pneumonia (PJP) in pediatric patients undergoing autologous stem cell transplantation (ASCT) during the period of pancytopenia prior to engraftment.

Our hospital is a pediatric referral center for childhood cancer. IVP prophylaxis for PJP in ASCT patients has been used since 2007. We did a retrospective review of our ASCT database to describe our experience on this topic.

Between March 2007 and January 2017, IVP for PJP prophylaxis was used in 92 ASCT carried out in 55 patients (36 male and 19 female). Of the 55 patients, 19 received two to four tandem ASCT as per disease protocol. The main indications for SCT were 39 solid tumors (29 central nervous system tumors, 8 neuroblastomas, and 2 Wilms tumors) and 16 hematological malignancies (12 acute myeloid leukemia, 2 acute lymphoblastic leukemia, and 2 lymphomas). The median age was 4.3 years (range 0.7–18.4). It is of note that six patients (10.9%) were younger than 2 years of age. There is some literature that suggests that infants under 2 years of age receiving prophylaxis with IVP may be at a higher risk of PJP infection than older children.<sup>3</sup>

The IVP dose was 4 mg/kg (maximum 300 mg) infused over 1 hr, every 2–3 weeks. The first dose coincided with the administration of the conditioning for SCT. The mean number of doses received per patient was 4 (range 2–11) for those receiving one ASCT and 6 (range 3–14) for tandem ASCT. The main reason for discontinuing IVP prophylaxis was switching the patients to the first-line treatment with trimethoprim–sulfamethoxazole (TMP–SMX) due to hematologic recovery. In our review, we did not find any breakthrough of PJP.

Only three patients (5.4%) had infusion-related adverse reactions. Two of them were serious reactions (anaphylaxis and seizures), leading to treatment discontinuation. One patient experienced nausea and hypotension that resolved, slowing the infusion rate. The incidence of adverse events was comparable to other published data on safety of IVP prophylaxis.<sup>2–5</sup>

While TMP–SMX remains the gold standard for PJP prophylaxis,<sup>7</sup> bone marrow toxicity is a limiting factor for its use in pediatric patients receiving tandem ASCT, who require adequate hematologic recovery between them. American and European guidelines suggest second-line drugs such as inhaled pentamidine, dapsone, or atovaquone, with

limited evidence.<sup>1,6,7</sup> Some practicalities should be considered, related to equipment and facilities that may make using inhaled pentamidine difficult.<sup>2,6</sup>

Our series of ACST pediatric patients treated with IVP is homogeneous in indication and dosage regimen. In our experience, prophylaxis with IVP has proved to be safe, feasible, and effective, with a breakthrough rate of 0. In addition, no toxoplasmosis infections have been reported.

We hope our experience can increase the evidence on the use safety, feasibility, and efficacy of IVP as the second-line PJP prophylaxis when TMP–SMX is not suitable.

### CONFLICT OF INTEREST

The authors declare that there is no conflict of interest.

Catalina Montoya Tamayo  
 Department of Pediatric Hematology and Oncology, Hospital Sant Joan de Déu, Barcelona, Spain

Ferran Bossacomà Busquets and Joan Vinent Genestar  
 Department of Pharmacy, Hospital Sant Joan de Déu, Barcelona, Spain

Claudia Fortuny Guasch  
 Department of Pediatric Infectious diseases, Hospital Sant Joan de Déu, Barcelona, Spain

Vicente Santa-Maria López and Susana Rives  
 Department of Pediatric Hematology and Oncology, Hospital Sant Joan de Déu, Barcelona, Spain

### Correspondence

Catalina Montoya Tamayo, Department of Pediatric Hematology and Oncology, Hospital Sant Joan de Déu, Passeig Sant Joan de Déu, 2, Esplugues de Llobregat, Barcelona 08950, Spain.  
 Email: cmontoya@sjdhospitalbarcelona.org

### REFERENCES

1. Kruizinga MD, Bresters D, Smiers FJ, Lankester AC, Bredius RGM. The use of intravenous pentamidine for the prophylaxis of *Pneumocystis pneumonia* in pediatric patients. *Pediatr Blood Cancer*. 2017;00:e26453.
2. Solodokin LJ, Klejmont LM, Scipione MR, Dubrovskaya Y, Lighter-Fisher J, Papadopoulos J. Safety and effectiveness of intravenous pentamidine for prophylaxis of *pneumocystis jirovecii* pneumonia in pediatric hematology/oncology patients. *J Pediatr Hematol Oncol*. 2016;38(6):e180–e185.

3. Clark A, Hemmelgarn T, Danziger-Isakov L, Teusink A. Intravenous pentamidine for *Pneumocystis carinii/jiroveci* pneumonia (PCP) prophylaxis in pediatric transplant patients. *Pediatr Transplant*. 2015;19:326–331.
4. DeMasi JM, Cox JA, Leonard D, Koh AY, Aquino JM. Intravenous pentamidine is safe and effective as primary pneumocystis pneumonia prophylaxis in children and adolescents undergoing hematopoietic stem cell transplantation. *Pediatr Infect Dis J*. 2013;32:933–936.
5. Kim SY, Dabb AA, Glenn DJ, Snyder KM, Chuk MK, Loeb DM. Intravenous pentamidine is effective as second line *Pneumocystis pneumonia* prophylaxis in pediatric oncology patients. *Pediatr Blood Cancer*. 2008;50(4):779–783.
6. Lim MJ, Stebbings A, Lim SJ, et al. IV pentamidine for primary PJP prophylaxis in adults undergoing allogeneic hematopoietic progenitor cell transplant [letter to the editor]. *Bone Marrow Transpl*. 2015;50:1253–1255.
7. Maertens J, Cesaro S, Maschmeyer G, et al. ECIL guidelines for preventing *Pneumocystis jirovecii* pneumonia in patients with haematological malignancies and stem cell transplant recipients. *J Antimicrob Chemother*. 2016;71(9):2397–2404.