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May 9th, 11:30 AM - 1:30 PM

A Novel Therapy for Refractory CBFA2T3::GLIS2-associated AMKL Using STRO-002 And Plerixafor

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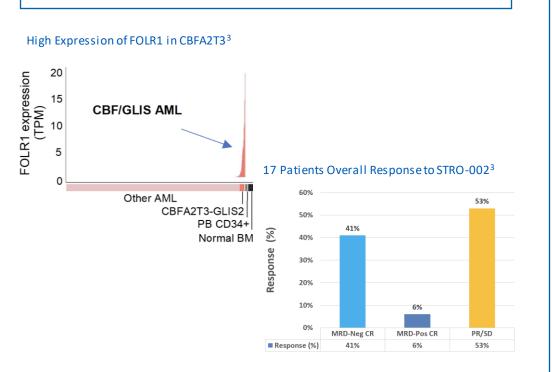
A Novel Therapy for Refractory CBFA2T3::GLIS2-associated AMKL Using **STRO-002 And Plerixafor**

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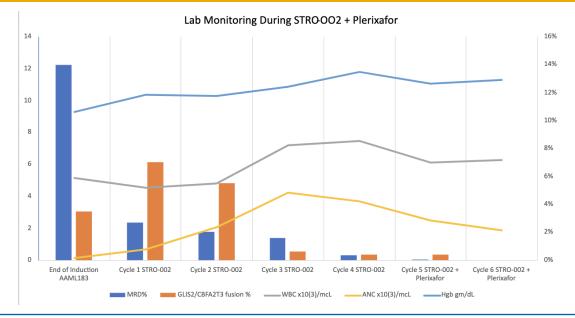
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Background:

- CBFA2T3::GLIS2-associated AML is an aggressive form of AML with a very poor prognosis with marrow localization of leukemic blasts likely mediated by high CXCR4 (CD184) expression.¹
- The CBFA2T3::GLIS2 fusion gene is among the most common oncogenic transcript in pediatric AMKL with high expression of CXCR4. Plerixafor, a reversible CXCR4 antagonist, mobilizes marrow stem cells and leukemic cells.²
- STRO-002 is an antibody/drug conjugate targeting FOLR1. A summary of 17 patients with refractory CBRA2T3::GLIS2 AML who received STRO-002 reported significant clinical activity with little to no toxicity.³





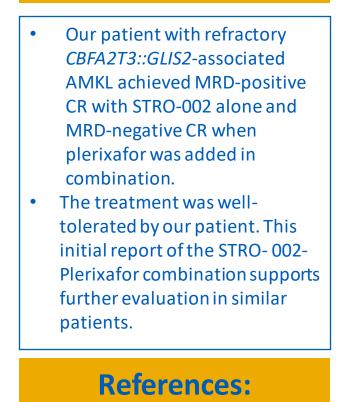


- A 2-year-old female presented with fever, arm pain, and bruising and was diagnosed with RAM phenotype AMKL with FOLR1-positive CBFA2T3::GLIS2 oncogenic fusion.
- She was enrolled on AAML1831 Arm A but Induction therapy was unsuccessful.
- She was transitioned to STRO-002 monotherapy (4.3mg/kg/dose IV every 2 weeks) as an outpatient for 4 cycles. Bone Marrow was assessed every 2 weeks.
- Plerixafor was added for leukemic cell mobilization with Cycles 5 and 6 (Plerixafor 0.24mg/kg/dose 4h prior and 24h post each STRO-002 dose).
- After Cycle 6, our patient achieved 0% MRD by flow and 0% CBFA2T3::GLIS2 fusion expression.
- She underwent haploidentical bone marrow transplant, but unfortunately relapsed on Day 100.
- Plan to continue STRO-002 with Plerixafor and Donor Leukocyte Infusions.





Conclusion:



1.	Andrew J. Menssen, Chad A. Hudson, Todd A. Alonzo, Robert B. Gerbing, Laura Pardo, Fan-Chi Hsu, Loren L. Lott, Fangyan Dai, Keely Ghiradelli, Yi-Cheng Wang, E. Anders Kolb, Todd M. Cooper, Jessica A. Pollard, Michael R. Loken, Richard Aplenc, Lisa Eidenschink Brodersen, Soheil Meshinchi; CXCR4 (CD184) Expression in Pediatric AML Is Associated with Bone Marrow Retention, Specific Disease Characteristics, and Worse Outcomes: A Report of 1004 Patients from the Children's Oncology Group AAML1031 Protocol. <i>Blood</i> 2022; 140 (Supplement 1): 3452–3453. doi: https://doi.org/10.1182/blood-2022-163682
2.	Tang T, Le Q, Castro S, Pardo L, McKay CN, Perkins L, Smith J, Kirkey D, Abrahams C, Bedard K, Molina A, Brodersen LE, Loken MR, Tarlock K, Meshinchi S, Loeb KR. Targeting FOLR1 in high-risk CBF2AT3-GLIS2 pediatric AML with STRO-002 FOLR1-antibody-drug conjugate. Blood Adv. 2022 Nov 22;6(22):5933-5937. doi: 10.1182/bloodadvances.2022008503. PMID: 36149945; PMCID: PMC9701621.
3.	Meshinchi, S. (2022, December 10-13). Anti-Leukemic Activity of STRO-002, a Novel Folate Receptor-α (FR-α)- Targeting ADC in Relapsed/Refractory CBFA2T3::GLIS2 AML. 64th ASH Annual Meeting and Exposition.

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