



Navigating Rett Syndrome Therapies: Essentials for Pediatric Care¹⁻¹³

Full abbreviations, accreditation, and disclosure information available at [PeerView.com/Q8540](https://www.peerview.com/Q8540)

Review of Evidence for Trofinetide in Rett Syndrome

Trial	Design	Population	Key Findings
LAVENDER	12-week, phase 3, double-blind, placebo-controlled	<ul style="list-style-type: none"> Females 5-20 years of age Classic RTT Confirmed MECP2 mutation 	<ul style="list-style-type: none"> Statistically significant improvement in global measures of function Favorable safety profile with mostly mild GI AEs Diarrhea; self-limiting, resolved after withdrawal of trofinetide
LILAC	Open-label extension of LAVENDER through 40 weeks	LAVENDER population	<ul style="list-style-type: none"> Sustained improvements in symptoms Consistent safety with no new concerns
DAFFODIL	78-week, open-label, phase 2/3	<ul style="list-style-type: none"> Females 2-5 years of age Classic RTT Confirmed MECP2 mutation 	<ul style="list-style-type: none"> Confirmed efficacy in global pediatric population Reduced frequency of adverse events, with diarrhea most common but manageable

Overview of Emerging Therapies

Agent	Mechanism of Action	Administration	Trial Status
Bilacamesine	Sigma-1 receptor agonist and muscarinic receptor modulator	Oral administration, once daily	<ul style="list-style-type: none"> Phase 3 trial in women 18-45 years: 72.2% of treatment arm vs 38.5% of placebo arm showed a significant improvement in RSBQ Phase 2/3 EXCELLENCE trial in children ≥5-17 years: significant improvements in RSBQ scores at week 4; favorable trends at week 12
TSHA-102	AAV9-mediated delivery of miniMECP2 gene	Single intrathecal injection	<ul style="list-style-type: none"> Currently in phase 2 (NCT05655566), moving into phase 3
NGN-401	AAV9-mediated MECP2 gene therapy	Single intracerebroventricular administration	<ul style="list-style-type: none"> Currently in phase 1/2 (REVEAL series), moving into phase 3

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