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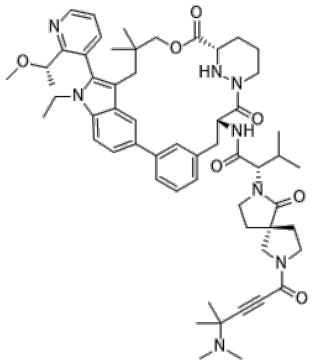
Data Sheet

Global Supplier of Chemical Probes, Inhibitors & Agonists.

Product Name	:	RMC-4998
Cat. No.	:	PC-21063
CAS No.	:	2642037-07-6
Molecular Formula	:	$C_{57}H_{74}N_8O_7$
Molecular Weight	:	983.25
Target	:	Ras
Solubility	:	10 mM in DMSO



Biological Activity

RMC-4998 (RM-029) is a tricomplex inhibitor that targets the active state or GTP-bound state of **KRAS G12C**, selectively inhibits the proliferation of KRASG12C mutant cells with mean IC50 of 0.28 nM.
RMC-4998 exhibits greater binding affinity (inhibition constant (KI): 272,000 M−1 s−1 (kinact/KI): 272,000 M−1s−1]) than that of existing inactive state-selective inhibitors.
RMC-4998 induces a tricomplex between KRASG12C and CYPA in live cells without affecting wild-type KRAS, NRAS, or HRAS.
RMC-4998 shows comparable activity on G12C-mutant HRAS, NRAS, and KRAS, with little, if any, activity on KRASG13C.
RMC-4998 (0.1 µM) led to a faster disruption of the interaction between KRASG12C and CRAF compared with inactive state-selective inhibitors adagrasib (1 µM) and sotorasib (10 µM) in mutant KRAS in cancer cells.
RMC-4998 inhibits KRASG12C-driven tumor growth in vivo.

References

Christopher J Schulze, et al. ***Science.*** 2023 Aug 18;381(6659):794-799.

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