

(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property
Organization
International Bureau

(43) International Publication Date
17 April 2025 (17.04.2025)



(10) International Publication Number
WO 2025/080653 A1

(51) International Patent Classification:

C07D 487/04 (2006.01) A61K 31/53 (2006.01)
C07D 491/04 (2006.01) A61K 35/00 (2006.01)
C07D 498/04 (2006.01)

(21) International Application Number:

PCT/US2024/050482

(22) International Filing Date:

09 October 2024 (09.10.2024)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data:

63/589,174 10 October 2023 (10.10.2023) US

(71) Applicant: **SHY THERAPEUTICS, LLC** [US/US]; 34
Bates Road, Harrison, NY 10528 (US).

(72) Inventors: **HADARI, Yaron, R.**; 34 Bates Road, Harrison, NY 10528 (US). **SCHMERTZLER, Michael**; 2270 Pinellas Point Dr. S., St. Petersburg, FL 33712 (US). **WILLIAMS, Theresa, M.**; 260 Shady Nook Poad, Harleysville, PA 19438 (US). **BIGARELLA, Carolina**; 37201 Town Green Drive, Elmsford, NY 10523 (US). **CARTA, Luca**; 2 Fountain Ln #3H, Scarsdale, NY 10583 (US). **HUTCHESON, Rebecca**; 389 Courtland Ave, Apt 1A, Stamford, CT 06906 (US). **ZHANG, Sufang**; 215 Schrade Road, Apt 2C, Briarcliff Manor, NY 10510 (US).

(74) Agent: **KASSAB, Darren, J.** et al.; Jones Day, 250 Vesey Street, New York, NY 10281-1047 (US).

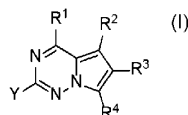
(81) Designated States (unless otherwise indicated, for every kind of national protection available): AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BN, BR, BW, BY, BZ, CA, CH, CL, CN, CO, CR, CU, CV, CZ, DE, DJ, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IQ, IR, IS, IT, JM, JO, JP, KE, KG, KH, KN, KP, KR, KW, KZ, LA, LC, LK, LR, LS, LU, LY, MA, MD, MG, MK, MN, MU, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PA, PE, PG, PH, PL, PT, QA, RO, RS, RU, RW, SA, SC, SD, SE, SG, SK, SL, ST, SV, SY, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, WS, ZA, ZM, ZW.

(84) Designated States (unless otherwise indicated, for every kind of regional protection available): ARIPO (BW, CV, GH, GM, KE, LR, LS, MW, MZ, NA, RW, SC, SD, SL, ST, SZ, TZ, UG, ZM, ZW), Eurasian (AM, AZ, BY, KG, KZ, RU, TJ, TM), European (AL, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, ME, MK, MT, NL, NO, PL, PT, RO, RS, SE, SI, SK, SM, TR), OAPI (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, KM, ML, MR, NE, SN, TD, TG).

Published:

— with international search report (Art. 21(3))

(54) Title: PYRROLO[2,1-F][1,2,4]TRIAZINE COMPOUNDS ACTING AGAINST CANCERS, INFLAMMATORY DISEASES, AND FIBROTIC DISEASE VIA INTERACTION WITH RAS SUPERFAMILY PROTEINS



(57) Abstract: Provided herein are compounds of Formula (I), or a pharmaceutically acceptable form thereof, and pharmaceutical compositions comprising the same. Also provided herein methods of modulating the activity of cellular targets by administering to a subject a compound of Formula (I), or a pharmaceutically acceptable form thereof. Further provided herein are methods of treating cancer, fibrotic diseases, and inflammatory diseases by administering to a subject a compound of or a pharmaceutically acceptable form thereof.



WO 2025/080653 A1