
Kenichirou Yokota¹, Masayori Hagimori*², Naoko Mizuyama³, Yasuhisa Nishimura¹, Hiroshi Fujito³, Yasuhiro Shigemitsu⁴ and Yoshinori Tominaga*¹

Abstract
New fluorescent compounds, benzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxides (3a–g), 2-amino-4-methylsulfanylbenzo[4,5]thieno[3,2-d]pyrimidine (6), and 2-amino-4-methylsulfanyl-7-methoxybenzo[4,5]furo[3,2-d]pyrimidine (7), were synthesized in good yields from heterocyclic ketene dithioacetals (1a–c) and guanidine carbonate (2a) or (S)-methylisothiourea sulfate (2b) in pyridine under reflux. Among the fused pyrimidine derivatives, compound 3c, which has an amino group at the 2-position and a benzylamino group at the 4-position of the pyrimidine ring, showed the strongest solid-state fluorescence. The absorption and emission properties of the compounds were quantitatively reproduced by a series of ab initio quantum-chemical calculations.

Introduction
Solid-state fluorescent compounds are currently attracting considerable interest from both theoretical and practical standpoints [1-4]. Recently, we prepared new pyrimidine derivatives, which have solid-state blue fluorescence, by means of a one-pot synthesis, involving the reaction of ketene dithioacetals, amines, and guanidine carbonate in pyridine [5]. We have also reported
the one-pot synthesis of a new, fluorescent, fused pyrimidine derivative 2,4-diaminoindeno[1,2-d]pyrimidin-5-one; this pyrimidine derivative, which was synthesized by heating a ketene dithioacetal, 2-[bis(methylsulfanyl)methylidene]indan-1,3-dione, under reflux with amine and amidine derivatives in pyridine solution, showed blue-green fluorescence in the solid state [6]. These one-pot synthetic methods are also promising for the preparation of other new solid-state fluorescent pyrimidine derivatives containing polycyclic heterocycles. In this paper, we report the synthesis of new fluorescent 2-aminobenzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxides and related fused pyrimidine derivatives. It is thought that these derivatives show strong fluorescence as a result of the presence of a hetero-π-electron conjugated system [7]. The electronic and emission spectra of these new compounds were analyzed computationally by using a series of ab initio quantum-chemical calculations.

Results and Discussion

Synthesis

Heterocyclic ketene dithioacetals (1a–c) were easily prepared by the condensation of the corresponding heterocyclic active methylene compounds with carbon disulfide in sodium hydroxide solution as a base, followed by methylation with dimethyl sulfate [8-10]. The reaction of 1a with guanidine carbonate (2a) in pyridine under reflux gave the expected product, 2-amino-4-(methylsulfanyl)benzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxide (3a) in 89% yield (Scheme 1). Based on our findings that diaminopyrimidine derivatives show blue fluorescence in the solid state [5], we attempted to synthesize diaminopyrimidine derivatives (3b–e), expecting them to show intense solid-state fluorescence. 2,4-Diaminobenzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxide (3b) was easily obtained, in 47% yield, by the displacement reaction of 3a with 28% ammonia solution at 200 °C for 3 h in a mini-autoclave.

Compound 3b was also synthesized by the reaction of 2a with 2-(diaminomethylene)benzol[b]thiophene-3(2H)-one 1,1-dioxide (4), which was easily obtained from 1a by using 28% ammonia in methanol under reflux. The reactions of 1a, 2a, and amines 5a–c (benzylamine, piperidine, and aniline) in pyridine under reflux gave diaminopyrimidine derivatives 3c–e in good yields, i.e., 92%, 51%, and 86%, respectively. The reaction of 1a and 2a under reflux in methanol instead of pyridine gave the methoxy pyrimidine derivative 3f in 55% yield.

In the presence of sulfuric acid solution, 2,4-dimethylsulfanyl derivative 3g was obtained from 1a with (S)-methylisothiourea sulfate (2b) in 87% yield (Scheme 2). Fused pyrimidine derivatives 6 and 7 containing the electron-rich heterocycles benzothiophene or benzofuran were prepared by the reaction of ketene dithiaoacetal 1b or 1c with guanidine carbonate (2a) in 74% and 65% yield, respectively.

Computational methods
The ground-state (S\textsubscript{0}) geometries were optimized by means of density functional theory (DFT) with the B3LYP hybrid functional and 6-311G(d,p) [11] basis set (DFT(B3LYP)/6-311G(d,p)). The first excited-state (S\textsubscript{1}) geometries were optimized with configuration interaction singles CIS(6-311G(d,p)) and the complete active space SCF CASSCF(10,10)/ANO-S-MB [12] level of theory. Time-dependent TDF (TDDFT), CASSCF, and multistate complete active-space second-order perturbation theory (MS-CASPT2) calculations were subsequently performed for the optimized geometries to evaluate the vertical transition energies along with the associated transition characteristics. The TDDFT calculations consistently employed the B3LYP functional and 6-31+G(d,p) basis set [13]. Solvent effects were evaluated by using the linear-response polarized-continuum model for ethanol (PCM(ethanol)-TDDFT(B3LYP)/6-31G+(d,p)) [14]. MS-CASPT2 used ten active electrons distributed within eight active spaces combined with an ANO-S basis set (MS-CASPT2(10,8)/ANO-S). The DFT, CIS, and TDDFT computations were performed by using Gaussian 09 software [15], and CASSCF and MS-CASPT2 were performed by using the MOLCAS 7.4 suite of programs [16].

Comparison of predicted and experimental absorption spectra
The UV–vis spectra of the compounds exhibited peaks (λ\textsubscript{max}) at 280–360 nm. The spectra possessed multiple subpeaks around λ\textsubscript{max}. The experimental and computational λ\textsubscript{max} values are given in Table 1. The theoretical and experimental λ\textsubscript{max} values are in fairly good agreement, within 50 nm deviation. The redshifts observed for 3d, 3e, and 6 relative to 3b were well reproduced by the TDDFT computations. The sidebands around λ\textsubscript{max} were not reproduced by our TDDFT computations, which predicted single-peak maxima for all the compounds. The subpeaks are considered to be vibronically assisted absorptions, because the extended π-systems of the molecules have large degrees of vibrational freedom effectively coupled with their electronic states. As a representative example of the electronic structures, the HOMO and LUMO of 3a are depicted in Figure 1. The λ\textsubscript{max} was assigned as the HOMO–LUMO π–π* excitation (configuration weight = 0.697) with an oscillator strength of 0.096 at 335 nm. The next theoretical peak appeared

Scheme 2: Syntheses of 3g, 6, and 7.
at 290 nm, derived from (HOMO – 1) to LUMO excitations, considerably separated from the first peak. The HOMO and LUMO delocalize on the whole system, indicating that modest intramolecular electron transfer from the methylthio and amino groups on the pyrimidine ring (HOMO) to the phenyl moiety (LUMO) is expected upon $S_0 \rightarrow S_1$ transition. The transition character is also rationalized with the transition dipole moment directed along the long molecular axis. The electron density difference between HOMO and LUMO is shown in Figure 2; this reflects the balanced electron redistribution between the pyrimidine and phenyl moieties.

**Table 1:** Experimental and computed absorption maxima ($\lambda_{\text{max}}$) for 3a–g, 6, and 7.

<table>
<thead>
<tr>
<th>No.</th>
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<th>$\lambda_{\text{max}}$ (calc.)$^b$</th>
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<td>7</td>
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<td>328</td>
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$^a$measured in dichloromethane, excluding 3b (in ethanol).

$^b$TD-DFT(B3LYP)/6-31+G(d,p)//DFT(B3LYP)/6-311G(d,p).

**Figure 1:** (a) HOMO and (b) LUMO for 3a, computed with TD-DFT(B3LYP)/6-31+G(d,p). The pink (blue) lobes indicate a positive (negative) isocontour value of 0.03.

**Figure 2:** The distribution of the HOMO minus LUMO density for 3a, computed with TD-DFT(B3LYP)/6-31+G(d,p). The pink (blue) lobes indicate a positive (negative) isocontour value of 0.003, respectively.

**Solid-state fluorescence**

Tris(8-hydroxyquinolinato)aluminium (Alq$_3$) was used as the standard for fluorescence spectrum measurements [17]. We analyzed the solid-state fluorescence emission spectra of 3a–g, 6 and 7 at room temperature. The fluorescence maxima ($\lambda_{\text{em,max}}$) and relative fluorescence intensities (RI) of these compounds are listed in Table 2. The $\lambda_{\text{em,max}}$ values of 3a and 3c–g were in the range 430–462 nm. The fluorescence of the $N$-unsubstituted diamino compound 3b exhibited a large bathochromic shift, and green fluorescence was observed ($\lambda_{\text{em,max}} = 529$ nm). Compound 3a, which has an amino group at the 2-position in the pyrimidine ring, showed stronger fluorescence than did compound 3g, which has a methylsulfanyl group. This result was in agreement with the previous findings that an amino group at the 2-position of an indeno pyrimidine influences the fluorescence intensity [6]. 2,4-Diaminopyrimidine derivatives (3b–e) also showed stronger fluorescence than did 3g. Compound 3c, which has an amino group at the 2-position and a benzylamino group at the 4-position of the pyrimidine ring, showed the strongest fluorescence (RI = 1.95). This value was larger than those of indeno pyrimidine derivatives (RI = 0.01–0.73). We speculated that the effect of the $\pi$-$\pi$-electron conjugated system in the sulfonyl group of benzothienopyrimidine 5,5-dioxide is stronger than that of the carbonyl group of the indeno pyrimidines.

Benzo[4,5]thieno[3,2-\text{d}]pyrimidine derivative 6 and benzo[4,5]furo[3,2-\text{d}]pyrimidine derivative 7 showed different emission properties from those of 3a–g. Compound 6 exhibited a large bathochromic shift compared with that of the benzo[4,5]thieno[3,2-\text{d}]pyrimidine 5,5-dioxides 3a and 3c–g.
and relatively strong fluorescence. In contrast, the weakly fluorescent compound 7 exhibited a hypsochromic shift relative to 3a–g. These results presumably indicate that hetero atoms incorporated in the ring moieties strongly influence the solid-state-fluorescent properties.

Solid-state fluorescence is strongly influenced by intermolecular steric hindrance. In general, the fluorescence intensity can be enhanced by minimizing intermolecular interactions through the introduction of bulky substituents. Compound 3e, which has a bulky benzamino moiety shows the strongest fluorescence of these molecules (Figure 2). The emission from 3e with a bulky phenylamino moiety, however, is relatively weak. This contradiction reflects the complex emission mechanism, which is driven not only by intermolecular stacking effects but also by the electronic structures. The weak fluorescence of 7 is assumed to be dominated by the inductive effect of the methoxy group, rather than by stacking effects.

To elucidate the S1 nature of the compounds in question, we carried out ab initio molecular-orbital calculations, focused on 3a (Figure 3). The computed key bond lengths are shown in Table 3. Comparison of the S0 (DFT) and S1 (CIS) bond lengths shows that the S0→S1 (dominant HOMO→LUMO) transition is reflected in the bond-length variations. The C4–C5 bond, with a bonding lobe in S0 but an antibonding one in S1, is significantly lengthened, by 0.045 Å, as a result of the S0→S1 transition. In contrast, the C5–C8 bond, with an antibonding lobe in S0 but a bonding one in S1, is considerably shortened, by 0.081 Å, on S0→S1 excitation. No significant deviation from planarity was observed in the fused-ring structure associated with the S0→S1 excitation. The geometrical discrepancies between DFT and CAS for S0 and CIS and CAS for S1 indicate that the significant roles of dynamic electron correlations are inadequate in the CAS method, which consistently predicted longer bond lengths. These disagreements could be resolved by geometric optimizations using MS-CASPT2 including dynamic correlation, but this is not feasible in the present study owing to the huge computational burden.

<table>
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<sup>a</sup>Stokes shift; emission - excitation in solid state. <sup>b</sup>Relative intensity of fluorescence in solid states, using Alq3 as the standard compound.

Theoretical fluorescence λ<sub>max</sub> values of 3a, obtained using several quantum-chemical methods, are summarized in Table 4. The CIS-predicted peak shows a significant blue-shift relative to the experimental value; this is well known, and is a result of the lack of multi-excitation character. The TDDFT peaks partially improve the gap, with the CIS optimized geometry for the S1 state not being of trustworthy quality. The CASSCF peak, in contrast, overshoots the experimental peak. The best prediction was obtained by using the most elaborate method, MS-CASPT2, which can take the majority of the electronic correlations into account. The MS-CASPT2 result indicates that...
were obtained on Gemini 300NMR (300 MHz) and 500NMR spectrometers. Nuclear-magnetic-resonance (NMR) spectra were determined on a Shimazu RF-1500 spectrometer. Fluorescence spectra were determined on a Hitachi JASCO 810 or Shimazu IR-460 spectrometer. Ultraviolet (UV) absorption spectra were determined in 95% ethanol on a Hitachi UV-160 spectrometer. Infrared (IR) spectra were recorded in potassium bromide pellets on a Perkin-Elmer 577 spectrometer. Melting points were determined with a Mita-Mart apparatus and were uncorrected. Infrared spectra were measured with a Shimazu IR-460 spectrometer. Microanalyses were performed by H. Mazume on a Yanaco M-5 at Nagasaki University. All chemicals were reagent grade and used without further purification unless otherwise specified.

**Conclusion**

New solid-state fluorescent compounds, benzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxides (3a–g), were synthesized in good yields by a convenient one-pot reaction of 2-[bis(methylsulfonyl)methylene]benzo[b]thiophene-3(2H)-one 1,1-dioxide (1a) with guanidine carbonate (2a) or (S)-methylisothiourea sulfate (2b) under reflux in pyridine. Benzo[4,5]thieno[3,2-d]pyrimidine derivative 6 and benzo[4,5]furo[3,2-d]pyrimidine derivative 7 were also synthesized by one-pot reactions between 2a and 1b or 1c, under the same conditions. The products 6 and 7 showed solid-state fluorescence. Compound 3c, which has an amino group at the 2-position and a benzylamino group at the 4-position of the pyrimidine ring, showed the strongest solid-state fluorescence. These results indicated that heteroatoms in the ring moieties have a strong influence on the solid-state emissions. The absorption and emission spectra of the compounds were computationally analyzed by means of a series of ab initio quantum-chemical calculations. The theoretical analyses quantitatively reproduced the S0→S1 and S1→S0 transitions. The associated HOMO/LUMO distributions and the transition characters were also elucidated theoretically.

**Experimental**

**General**

Identifications of compounds and measurements of properties were carried out by general procedures employing the following equipment: All melting points were determined with a Mitamura Riken Kogyo Mel-Temp apparatus or a Laboratory Devices Mel-Temp II apparatus and were uncorrected. Infrared (IR) spectra were recorded in potassium bromide pellets on a JASCO 810 or Shimazu IR-460 spectrometer. Ultraviolet (UV) absorption spectra were determined in 95% ethanol on a Hitachi 323 spectrometer. Fluorescence spectra were determined on a Shimazu RF-1500. Nuclear-magnetic-resonance (NMR) spectra were obtained on Gemini 300NMR (300 MHz) and 500NMR (500 MHz) spectrometers with tetramethylsilane as an internal standard. Mass spectra (MS) were recorded on a JOEL DX-303 mass spectrometer. Microanalyses were performed by H. Mazume on a Yanaco M-5 at Nagasaki University. All chemicals were reagent grade and used without further purification unless otherwise specified.

**Method of measurement of fluorescence**

A powder sample of the subject compound was heaped in the tray. After the sample was covered with a quartz plate, this part was fixed in the fluorescence spectrometer. After fixing the fluorescence wavelength, the excitation spectrum was determined by scanning with the fluorescence wavelength. Similarly, the fluorescence spectrum was obtained after scanning with the excitation wavelength. After obtaining these results, the excitation wavelength was decided and the fluorescence spectrum was measured. The relative intensity of fluorescence was determined by using Alq3 as the standard sample. The fluorescence of the standard sample and all subject compounds was measured at 272 nm excitation.

**Synthesis**

2-Amino-4-(methylsulfonyl)benzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxide (3a): A mixture of 2-[bis(methylsulfonyl)methylene]benzo[b]thiophene-3(2H)-one 1,1-dioxide (1a; 1.43 g, 5.0 mmol), guanidine carbonate (2a; 0.90 g, 5.0 mmol) and pyridine (10 mL) was stirred for 5 h under reflux. The reaction mixture was poured into ice water (100 mL). The colorless crystals that appeared were collected by filtration and washed with methanol (30 mL) to give 3a (1.28 g, 89%) as colorless needles. An analytical sample was recrystallized from a mixture of toluene and methanol to give colorless needles. Mp 294–296 °C; IR (KBr, cm−1): ν: 3409, 3332, 3219, 1650, 1546, 1520, 1145; UV (CH2Cl2) λmax (log ε): 234 (4.15), 396 (3.99), 461 (3.77); UV (EtOH) λmax (nm (log ε): 252 (4.43), 297 (3.91), 307 (4.04), 366 (3.92), 376 (3.93); 1H NMR (300 MHz, DMSO-d6) δ 2.64; s, (3H, SMe); 7.86 (m, 2H, 7-H, 8-H); 7.95–8.04 (m, 4H, NH2, 6-H, 9-H); 13C NMR (100 MHz, DMSO-d6) δ 110.1, 114.4, 121.5, 122.8, 129.5, 134.3, 134.4, 140.3, 158.5, 164.1, 165.2; MS m/z (% relative intensity): 280 (M+ + 1, 17), 279 (M+, 100), 262 (29), 215 (36), 209 (25).
2-(Diaminomethylene)benzo[θ]thiophene-(3H)-one 1,1-dioxide (4): A mixture of 1a (1.43 g, 5.0 mmol), ammonia solution (28%, 5 mL) and methanol (10 mL) was heated under reflux for 30 min. After cooling, water and methanol were evaporated. The residue was collected, washed with water and methanol, and crystallized from methanol to give 0.98 g (4.4 mmol, 88%) of colorless needles. Mp 312–314 °C; MS m/z (% relative intensity): 225 (M+ + 1, 1), 224 (M+, 100), 159 (11), 153 (24), 104 (11), 76 (13). Anal. calc’d for C8H6N2O2S: C, 48.21; H, 3.60; N, 12.49; found: C, 48.27; H 3.42; N, 12.51.

2,4-Diaminobenzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxide (3b): Method A: A mixture of 0.30 g (1.2 mmol) of 3a and ammonium hydroxide (28%, 20 mL) in a mini-autoclave (50 mL) was heated at 200 °C for 3 h. After cooling, the precipitate that appeared was collected by filtration and washed with water to give pale yellow needles (0.14 g, 0.6 mmol, 47%). Mp 315–318 °C (dec.). Method B: A mixture of 4 (0.30 g, 1.2 mmol), 2a (0.50 g, 3.0 mmol) and pyridine (10 mL) was refluxed for 7 h. After cooling, the reaction mixture was poured into water (50 mL). The precipitate that appeared was collected by filtration to give pale yellow needles (0.12 g, 0.5 mmol, 41%). Mp 320–322 °C (dec.). An analytical sample was recrystallized from methanol to give pale yellow needles. Mp 320–322 °C (dec.). IR (KBr, cm⁻¹): 3400, 3464, 3329, 3213, 2942, 1633, 1558, 1593, 1283, 993, 551; UV (CH2Cl2) λmax, nm (log e): 274 (3.64), 397 (4.03); EI NMR (300 MHz, CDCl3) δ 4.06 (d, J = 5.6 Hz, 2H, N-CH2-), 5.31 (br s, 2H, NH2), 5.64 (br s, 1H, NH), 7.28–7.38 (m, 5H, phenyl-H), 7.68 (m, 2H, 7-H, 8-H), 7.82 (m, 1H, 9-H), 7.99 (m, 1H, 5-H); 13C NMR (100 MHz, DMSO-d6) δ 42.9, 101.2, 120.9, 122.3, 126.7, 127.5, 128.2, 130.7, 133.3, 133.8, 139.5, 140.3, 155.5, 159.3 165.3; MS m/z (% relative intensity): 339 (M+ + 1, 21), 338 (M+, 100), 321 (62), 304 (25), 303 (44), 201 (23), 91 (47). Anal. calc’d for C17H14N4O2S: C, 57.64; H, 2.57; N, 16.65; found: C, 56.85; H, 2.64; N, 16.51.

2-Amino-4-piperidinylbenzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxide (3d): This compound (0.32 g, 1.0 mmol) was prepared in 51% yield from 1a (0.50 g, 2.0 mmol), piperidine (5b; 0.34 g, 4.0 mmol), and 2a (0.30 g, 1.7 mmol) in a similar procedure to that described for the synthesis of 3c. An analytical sample was recrystallized from a mixture of toluene and methanol to give colorless needles. Mp 224–226 °C; IR (KBr, cm⁻¹): 3480, 3447, 3329, 3213, 2942, 1633, 1558, 1283, 993, 551; UV (CH2Cl2) λmax, nm (log e): 274 (4.16), 397 (4.03). EI NMR (300 MHz, CDCl3) δ 1.74 (m, 6H, -(CH2)3-), 3.91 (m, 4H, N-CH2-), 5.11 (br s, 2H, NH2), 7.66 (m, 2H, 7-H, 8-H), 7.81 (m, 1H, 9-H), 8.00 (m, 1H, 6-H); 13C NMR (100 MHz, DMSO-d6): 24.1, 25.7, 25.7, 46.8, 46.8, 102.2, 120.8, 122.3, 130.6, 13.5, 133.9, 138.6, 156.1, 160.9, 164.08; MS m/z (% relative intensity): 317 (M+ + 1, 10), 316 (M+, 53), 299 (100), 282 (23), 127 (12), 84 (11). Anal. calc’d for C12H16N2O2S: C, 56.94; H, 5.10; N, 17.71; found: C, 56.79; H, 5.11; N, 17.56.

2-Amino-4-phenylaminobenzo[4,5]thieno[3,2-d]pyrimidine 5,5-dioxide (3e): This compound (0.56 g, 1.7 mmol) was prepared in 86% yield from 1a (0.57 g, 2.0 mmol), aniline (5e; 0.33 g, 3.5 mmol), and 2a (0.30 g, 1.7 mmol) in a similar procedure to that described for the synthesis of 3c. An analytical sample was recrystallized from a mixture of toluene and methanol to give colorless needles. Mp 252–255 °C; IR (KBr, cm⁻¹): 3346 (NH), 3192 (NH), 1651, 1602, 1551, 1292, 1143; UV (CH2Cl2) λmax, nm (log e): 271 (4.65), 344 (4.15), 396 (3.99), 461 (3.77); EI NMR (500 MHz, CDCl3) δ 7.15 (m, 2H, phenyl-H), 7.24 (m, 2H, phenyl-H), 7.40 (br s, 1H, NH), 7.49 (br s, 1H, NH), 7.69 (d, J = 7.3 Hz, 2H, phenyl-H), 7.83–7.86 (m, 2H, 7-H, 8-H), 7.97–8.03 (m, 2H, 6-H, 9-H); 13C NMR (100 MHz, DMSO-d6) δ 102.2, 121.0, 122.3, 123.9, 124.0, 124.2, 128.2, 130.5, 133.6, 134.0, 138.2, 140.4, 154.3, 160.1, 165.1; MS m/z (% relative intensity): 326 (M+ + 2, 11), 325 (M+ + 1, 35), 324 (M+, 100), 323 (79), 260 (14), 259 (62), 91 (15). Anal. calc’d for C19H14N2O2S: C, 59.25; H, 3.73; N, 17.27; found: C, 59.58; H, 3.95; N, 17.25.
2.0 mmol), and methanol (10 mL) was refluxed for 1 h. After cooling, the precipitate that appeared was collected by filtration to give colorless crystals (0.29 g, 1.1 mmol) in 55% yield. An analytical sample was recrystallized from toluene to give colorless needles. Mp 298–302 °C; IR (KBr, cm⁻¹) v: 3476, 3399, 3343, 3222, 1654, 1551, 1374, 1301, 1158; UV (CH₂Cl₂) λ_max (nm (log ε)): 253 (4.75), 322 (4.10), 397 (4.00), 462 (3.79); ¹H NMR (300 MHz, CDCl₃) δ 4.04 (s, 3H, OMe), 7.83–7.88 (m, 2H, 7-H, 8-H), 7.94 (br s, 1H, NH), 7.95–8.03 (m, 2H, 6-H, 9-H), 8.04 (br s, 1H, NH); ¹³C NMR (100 MHz, DMSO-d₆) δ 54.2, 103.4, 121.4, 122.6, 129.8, 134.1, 134.4, 140.7, 151.5, 162.1; MS m/z (% relative intensity): 264 (M⁺ + 1, 15), 263 (M⁺, 100), 246 (12), 246 (12), 220 (18), 152 (16), 136 (15); Anal. calcld for C₁₁H₁₄N₂O₂S: C, 50.16; H, 3.45; N, 15.96; found: C, 50.18; H, 3.41; N, 15.91.

2,4-Bis(methylsulfanyl)benzo[4,5]thieno[3,2-d]pyrimidine (3g): This compound (0.54 g, 1.8 mmol) was obtained in 87% yield from 1a (0.57 g, 2.0 mmol), S methylsulfiniourethrae (2b; 0.56 g, 2.0 mmol), and pyridine (10 mL) in a manner similar to that described for the synthesis of 3a. An analytical sample was recrystallized from toluene to give colorless needles. Mp 233–234 °C; IR (KBr, cm⁻¹) v: 2929, 1513, 1461, 1306, 1153, 869; UV (CH₂Cl₂) λ_max (nm (log ε)): 253 (4.75), 322 (4.10), 397 (4.00), 462 (3.79); UV (EtOH) λ_max (nm (log ε)): 208 (3.94), 275 (4.28), 345 (3.35); ¹H NMR (300 MHz, CDCl₃) δ 2.68 (s, 3H, SMe), 2.74 (s, 3H, SMe), 7.74 (m, 2H, 7-H, 8-H), 7.88 (m, 1H, 9-H), 8.15 (m, 1H, 6-H); ¹³C NMR (100 MHz, CDCl₃) δ 117.1, 142, 121.0, 122.0, 123.8, 128.8, 135.0, 135.1, 155.2, 161.2, 176.7; MS m/z (% relative intensity): 311 (M⁺ + 1, 17), 310 (M⁺, 100), 295 (11), 293 (18), 249 (15), 246 (11), 173 (12), 136 (12), 114 (12); Anal. calcld for C₁₂H₁₀N₂O₂S₂: C, 46.43; H, 3.25; N, 9.02; found: C, 46.45; H, 3.20; N, 8.96.

2-Amino-4-(methylsulfanyl)benzo[4,5]thieno[3,2-d]pyrimidine (6): This compound (0.92 g, 3.7 mmol) was prepared in 74% yield from 1b (1.27 g, 5.0 mmol) and 2a (1.0 g, 5.6 mmol) in a manner similar to that described for the synthesis of 3a. An analytical sample was recrystallized from a mixture of toluene and methanol to give pale yellow needles. Mp 191–193 °C; IR (KBr, cm⁻¹) v: 3487 (NH), 3313 (NH), 3186 (NH), 1630, 1524, 1575, 722; UV (CH₂Cl₂) λ_max (nm (log ε)): 249 (4.79), 294 (4.37), 304 (4.42), 361 (4.34), 372 (4.34), 462 (3.79); ¹H NMR (300 MHz, CDCl₃) δ 2.69 (s, 3H, SMe), 6.88 (br s, 2H, NH₂), 7.55 (m, 1H, 8-H), 7.67 (m, 1H, 7-H), 8.06 (d, J = 7.8 Hz, 1H, 9-H), 8.19 (d, J = 8.3 Hz, 1H, 6-H); ¹³C NMR (100 MHz, CDCl₃) δ 11.5, 115.8, 123.4, 123.9, 125.2, 130.1, 133.0, 140.2, 156.8, 161.5, 163.1; MS m/z (% relative intensity): 249 (M⁺ + 2, 45), 248 (M⁺ + 1, 75), 247 (M⁺, 100), 246 (75), 202 (39), 201 (54), 200 (33), 187 (39), 146 (54), 114 (21); Anal. calcld for C₁₁H₁₄N₂S₂: C, 53.42; H, 3.67; N, 16.99; found: C, 53.56; H, 3.64; N, 16.93.

References


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