**Supplementary Material**

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| Table S1: Characteristics of All Subjects Tested For CDI Using Two-Step Algorithm From 2018-2022 |
| Demographic Characteristic  | Total(N=3414)[col. %] | NegativeN=[row %] | DiscordantN=[row %] | PositiveN=[row %] | Not IncludedN=[row %] |
| Age  |  |  |  |  |  |
| *<65*  | 2060[60.34] | 1433[61.50] | 318[62.11] | 80[50.31] | 229[55.45] |
| *≥65* | 1354[39.66] | 897[38.50] | 194[37.89] | 79[49.69] | 184[44.55] |
| Sex  |  |  |  |  |  |
| *Male*  | 1671[48.97] | 1178[50.58] | 222[43.36] | 82[51.57] | 189[45.87] |
| *Female*  | 1743[51.03] | 1151[49.42] | 290[56.64] | 77[48.43] | 223[54.13] |
| Race |  |  |  |  |  |
| *Amer. Indian/ Alaska Native* | 1[0.03] | 0[0] | 0[0] | 0[0] | 1[0.24] |
| *Asian*  | 50[1.47] | 36[1.55] | 7[1.37] | 2[1.26] | 5[1.22] |
| *Black Or African American*  | 2143[62.84] | 1432[61.48] | 359[70.12] | 98[61.64] | 254[61.95] |
| *Multiple*  | 39[1.14] | 28[1.20] | 6[1.17] | 3[1.89] | 2[0.49] |
| *White* | 846[24.81] | 594[25.50] | 105[20.51] | 37[23.27] | 110[26.83] |
| *Unknown* | 331[9.71] | 239[10.26] | 35[6.83] | 19[11.95] | 38[9.27] |
| Ethnicity |  |  |  |  |  |
| *Non-Hispanic /Latino*  | 3048[89.34] | 2060[88.45] | 474[92.58] | 137[86.16] | 377[91.73] |
| *Hispanic/ Latino* | 162[4.75] | 131[5.62] | 15[2.93] | 9[5.66] | 7[1.70] |
| *Unknown* | 198[5.81] | 138[5.92] | 23[4.50] | 13[8.18] | 24[5.84] |
| Missing Attribute Values: Sex N=2, Race N=4, Ethnicity N=3.  |

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| **Table S2: Distribution of Antibiotic Treatment Outcome (N=3414)** |
|  | TotalN [col %] | Treated[row %] | Untreated[row %] |
| Negative  | 2330[68.25] | 183[7.85] | 2147[92.15] |
| PCR**-**PCR**-**Ag**-**T**-** |
| Discordant  | All512 [15.00]*PCR+ Disc.*450 *PCR- Disc.*62 | All168 [32.81]*PCR+ Disc.*127*PCR- Disc*41 | All344 [67.19]*PCR+ Disc.*323*PCR- Disc*21 |
| PCR**+**Ag**+**T**-**PCR**+**Ag**-**T**+**PCR**+** Ag**-**T**-**PCR- Ag+T+PCR**-** Ag**+**T**-**PCR**-** Ag**-**T**+** |
| Positive  | 159[4.66] | 134[84.28] | 25[15.72] |
| PCR**+**Ag**+**T**+** |
| Missing PCR or Ag/T Test Result | 386[11.31] | 200[51.81] | 186[48.19] |
| Other/Unknown | 27[0.78] |  |  |
| Abbreviations: PCR=Polymerase Chain Reaction, Ag= Antigen, T=Toxin, %=percent of that testing category  |