

Table S1. Sample and culture distribution.

Sample	Total patients/ samples	Direct microscopy positive	Culture positive	Repeat isolation
Blood	235/ 449	Not applicable	1 (<i>A.nidulans</i>)	1 (<i>A.nidulans</i>)
Sputum	188/ 197	21	20 (12 <i>A. flavus</i> , 8 <i>A.fumigatus</i>)	6 (4 <i>A.flavus</i> , 2 <i>A.fumigatus</i>)
BAL	24/ 41	7	7 (1 <i>A.flavus</i> , 4 <i>A.fumigatus</i> , 2 <i>A.terreus</i>)	6 (1 <i>A.flavus</i> , 4 <i>A.fumigatus</i> , 1 <i>A.terreus</i>)
ETA	23/ 39	5	5 (2 <i>A.flavus</i> , 2 <i>A.fumigatus</i> , 1 <i>A.niger</i>)	5 (2 <i>A.flavus</i> , 2 <i>A.fumigatus</i> , 1 <i>A.niger</i>)
Pleural effusion	5/ 5	1	1 (1 <i>A.fumigatus</i>) (similar isolation from patient's sputum sample)	
Total	235/ 731	33	34 (9 <i>A.flavus</i> , 10 <i>A.fumigatus</i> , 2 <i>A.terreus</i> , 1 <i>A.niger</i> , 1 <i>A.nidulans</i>) (1 patient had same isolate with two samples)	18

[Note: BAL: bronchoalveolar lavage; ETA: endotracheal aspirate]

Table S2. Summary of few studies depicting hot-spot and other mutations in *cyp51A* gene of *A. fumigatus* imparting resistance to azole drugs.

Total no. of <i>A. fumigatus</i> , source and drug resistant	Mutations in <i>cyp51A</i>	No of isolates showing mutations	Breakpoints followed	Methodology and Endpoint	Country/ Region	Reference
12(clinical): ITRA	Codon 54: G54E, G54V, G54R; Codon 236: M236T, M236V, M236K	6	≤0.5µg/ml susceptible and >8 resistant	CLSI NCCLS Endpoint: Complete inhibition of growth	Madrid, Spain/ Europe	Diaz-Guerra Tm et al., 2003

130(clinical): ITRA	TR34, L98H; S297T, F495I; G54E/W, G138C, M220R/I/V/K	105; 12; 8	itraconazole and voriconazole: <2 mg/l (susceptible), 2mg/l (intermediate), >2 mg/l (resistant); posaconazole <0.25, 0.5 and >0.5 mg/l	Agar screening: 4/8 mg/l itraconazole in SDA CLSI M38-A	Nijmegen, Netherlands/ Europe	Snelders E et al., 2010
121 (clinical) ITRA, VOR, and POS	TR34/L98H	5	resistant: >2mg/l, itraconazole: and voriconazole and >0.25mg/l, posaconazole	EUCAST	Poland/ Europe	Nawrot U et al., 2017
3(environmental): Itraconazole 1: VOR 1: POS	TR34, L98H	4	susceptible for itraconazole and voriconazole≤1mg/l; posaconazole≤0.25 mg/l	Agar screening: 4 mg/l itraconazole in SDA EUCAST Endpoint: No visible growth	Copenhagen, Denmark/ Europe	Mortensen KL et al., 2010
1(clinical):Pan azole	G432S	1	resistance cut-offs for itraconazole and voriconazole: >2mg/l; posaconazole >0.25mg/l	ETest	Paris, France/ Europe	Alanio A et al., 2011
14(clinical): ITRA	TR34, L98H	14	Not applicable	CLSI M38-A	Nijmegen, Netherlands/ Europe	Mellado E et al., 2007
17(clinical): ITRA	TR34/L98H M220I G54W F219C	6 1 1 1	resistance cut-offs for itraconazole and voriconazole: >2mg/l; posaconazole >0.25mg/l	EUCAST Endpoint: No visible growth	Germany/ Europe	Bader O et al., 2013
1(clinical): ITRA and VOR	TR46/Y121F/ T289A	1	Not applicable	EUCAST	France/ Europe	Lavergne R-A et al., 2015
1(clinical): ITRA and VOR	TR46/Y121F/ T289A	1	Not applicable	Sensititre YeastOne YO10; CLSI M38-A2	Belgium/ Europe	Montesinos I et al., 2014
4(clinical): Pan azole	TR46/Y121F/T298A ; M172I, G448S M220R TR34/L98H	1 1 1	resistance cut-offs for itraconazole and voriconazole: >2mg/l; posaconazole >0.25mg/l	EUCAST	Nijmegen, Netherlands/ Europe	van Ingen J et al., 2015

21(clinical): ITRA and VOR	TR46/Y121F/ T289A TR34/L98H M172I/G448S S297T/F495I	3 15 1 2	voriconazole: resistant: >2mg/l, susceptible: ≤1 mg/l. itraconazole: resistant: >2mg/l, susceptible: ≤1 mg/l. posaconazole: resistant: >0.25mg/l, susceptible: ≤0.12 mg/l.	EUCAST	Nijmegen, Netherlands/ Europe	Fuhren J et al., 2015
0 (clinical/hospital environment)	N479D;F165L;N248 K;D262Y;M172V;F 46Y;E427K;D255E; N248T; Other silent mutations	9	resistant: ≥4 mg/l susceptible:<1 mg/l	CLSI M38-A2	Madrid, Spain/ Europe	Escribano P et al., 2011
220 (clinical) ITRA, VOR, and POS	M220V,TR46/Y121 F/T289A, G54R, G138S, TR34/L98H, G448S, F219S, M220I, G54E, G138C	26	resistant: voriconazole and itraconazole ≥4 mg/l; posaconazole ≥0.5 mg/l	CLSI M38-A2	USA/ North America	Wiederhold NP et al., 2016
0	N248K; I242V; A9T; A9T	5	resistance cut-offs for itraconazole and voriconazole: >2mg/l	Sensititre YeastOne YO10; CLSI M38-A2 Endpoint: Complete inhibition of growth	Toronto, Canada/ North America	Shalhoub S et al., 2015
38 (environmental)	TR46/Y121F/T289A , TR53, TR34/L98H	17	Not applicable	CLSI M38-A2	Bogotá, Colombia/ Latin America	Le Pape P et al., 2016
1 (clinical) ITRA	G54E	1 (fungal keratitis case)	Not applicable	CLSI M38-A2	Argentina/ South America	Leonardelli F et al., 2017
4(clinical): ITRA	M220I; G54R	1; 3	Not applicable	E-Test CLSI M38-A Endpoint: No visible growth	Beijing, China/ Asia	Chen J et al., 2005
38 (clinical) ITRA, VOR, and POS	TR34/L98H	3	resistant: >1, >1 or >0.5 mg/l to itraconazole, voriconazole and posaconazole, respectively	Sensititre YeastOne, CLSI M38-A2	Taiwan/ Asia	Wu CJ et al., 2015

50 (environmental), 16 (clinical) ITRA, VOR, and POS	TR34/L98H	1 (environmental), 2 (clinical)	resistant: ≥ 2 , ≥ 0.5 and ≥ 2 mg/l to itraconazole, posaconazole and voriconazole, respectively	E-test, CLSI M38-A2	Kuwait/ Middle East	Ahmad S et al., 2015
746 (clinical) ITRA, VOR, and POS	TR34/L98H	66 mutations detected; 10 undetected	Not applicable	CLSI M38-A	Turkey/ Middle East	Özmerdiven GE et al., 2015
418 (clinical) ITRA, VOR, and POS	TR34/L98H, G54R, F46Y, Y431S and G448S	6 mutations detected; 3 undetected	resistant: >2 mg/l, itraconazole: and voriconazole and >0.25 mg/l, posaconazole	CLSI M38-A2	Australia/ Oceania	Kidd SE et al., 2015
685 (clinical) ITRA, VOR, ISA and POS	(10) TR34/L98H, (1) G54E and one isolate with non-synonymous mutation	12	itraconazole, 1 μ g/ml; voriconazole, 1 μ g/ml; posaconazole, 0.5 μ g/ml; isavuconazole, 1 μ g/ml; amphotericin B, 4 μ g/ml and caspofungin, 0.25 μ g/ml	CLSI M38-A2	Delhi, India/ Asia	Chowdhary A et al., 2015

[NOTE: ITR: itraconazole; VOR: voriconazole; POS; posaconazole; ISA: isavuconazole; CLSI: Clinical and Laboratory Standards Institute; EUCAST: European Committee on Antimicrobial Susceptibility Testing]