

MALDI-TOF MS

Identification of dermatophytes



- Dermatophytes
- Overview literature
- Routine use MALDI-TOF MS in dermatophyte identification

Dermatophytes



- Invasion and infection of keratinized tissues
- Three anamorphic genera: *Epidermophyton*, *Microsporum* and *Trichophyton*
- Geophilic, zoophilic and anthropophilic species (natural habitat)
- Treatment cost – half a billion dollars annually USA^(*)
- Resistance to treatment is uncommon
- Proper identification: critical for treatment regime and detection of inoculum source

*Graser, Y., J. Scott, and R. Summerbell. 2008. The new species concept in dermatophytes-a polyphasic approach. Mycopathologia 166:239-256.

Classical identification (ID)



- Correlation of clinical manifestations of infection and observation of macro- and microscopic properties
 - experienced technologists, morphological characteristics
 - sterile mycelium (antifungal treatment)
- DNA sequence-based ID (gold standard)
 - expensive and time-consuming

**Both restricted by slow growth of dermatophytes
(up to 3 weeks)**

MALDI-TOF MS identification



ID dermatophytes:

- Improve accuracy
- Decrease analysis time



Overview literature

DOI:10.1111/ijms.16012
Medical Int. J. Mol. Sci. 2014, 15, 16012–16024; doi:10.3390/ijms150916012

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Mycopathologia (2015) 180:165–171
DOI 10.1007/s11046-015-9898-x



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Evaluation of the Bruker Matrix-Assisted Laser Desorption–Ionization Time-of-Flight Mass Spectrometry (MALDI-TOF MS) System for the Identification of Clinically Important Dermatophyte Species

Nilgün Karabıçak · Onur Karatuna · Macit İlkit ·
Işın Akyar

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Published: 11 September 2014

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Technical challenges for routine use

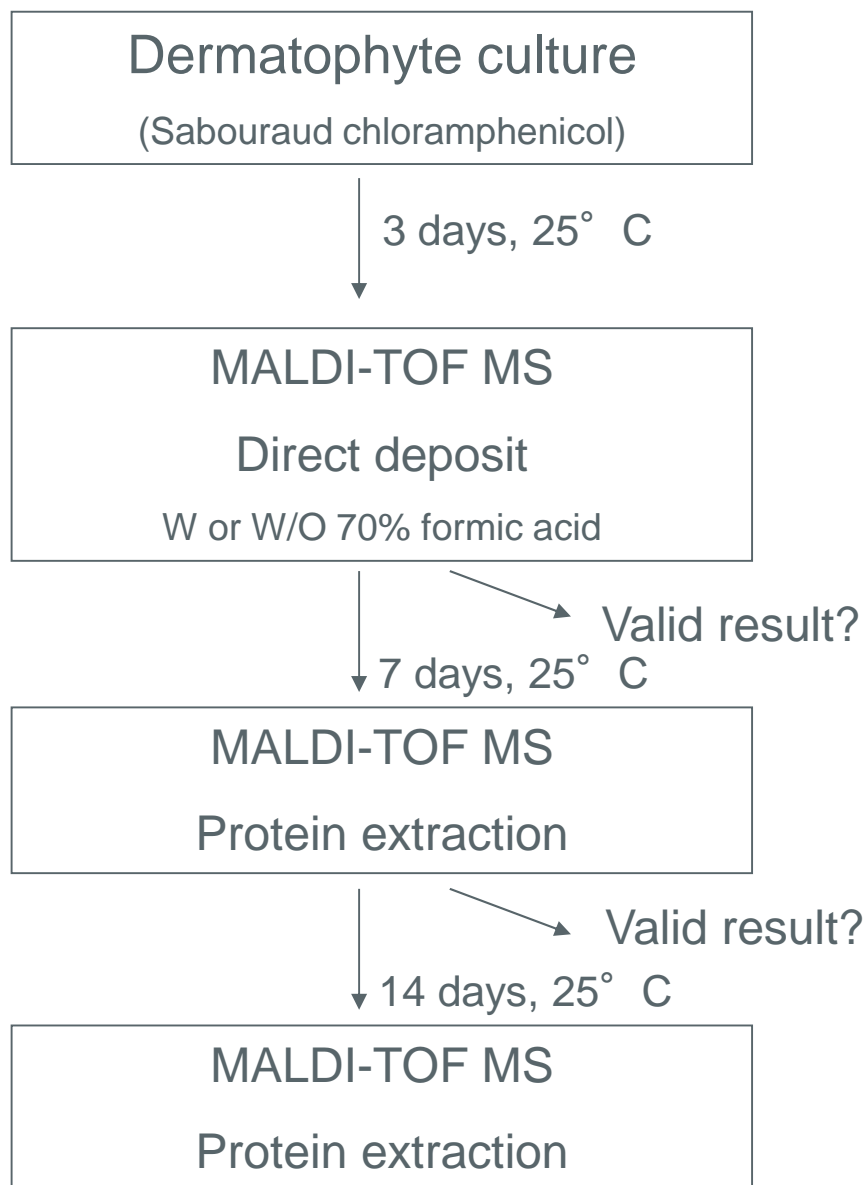


- Growth forms in sample (mycelium or conidia)
- Duration of incubation
- Culture medium
- Preparation methods (solid or liquid)
- Amount of biomass
- Type of instrument used

MALDI-TOF MS system	Author, year	Method	Result
Andromas	Alshawwa <i>et al.</i> , 2012	<ul style="list-style-type: none"> ref. db – 12 sp. 50 isolates 3w inc. direct + FA 	91.9% correct ID
AnagnosTec SARAMIS software	Erhard <i>et al.</i> , 2007	<ul style="list-style-type: none"> ref. db – 18 sp. 4w inc. direct 	99.9% correct ID (not: closely related sp.)
	Nenoff <i>et al.</i> , 2013	<ul style="list-style-type: none"> ref. db – 21 sp. 285 isolates at least 1w inc. direct 	98.8% correct ID (not: closely related sp.)
	De Respinis <i>et al.</i> , 2013	<ul style="list-style-type: none"> ref. db – 18 sp. 108 isolates 3d inc. direct + FA with sample prep 	95.8% correct ID (<i>T. rubrum</i> complex)
Vitek MS Plus System	De Respinis <i>et al.</i> , 2015	<ul style="list-style-type: none"> db + extension – 17 sp. 134 isolates 3d inc. in house sample prep. 	95.4% correct ID (<i>T. rubrum</i> complex)

MALDI-TOF MS system	Author, year	Method	Result
MALDI Biotyper (Bruker)	Theel <i>et al.</i> , 2011	<ul style="list-style-type: none"> db + extension – 20 spectra 3d inc. extr. 	20.5 vs 59.6% correct species ID
	Packeu <i>et al.</i> , 2013	<ul style="list-style-type: none"> ref. db – 54 isolates 12d inc. extr. 	<i>T. menta</i> complex – 89% correct ID
	l'Ollivier <i>et al.</i> , 2013	<ul style="list-style-type: none"> ref. db – 17 sp. 48 isolates 3/6d inc. extr. 	97.8% correct ID
	Packeu <i>et al.</i> , 2014	<ul style="list-style-type: none"> ref. db – 58 sp. 195 isolates 3/14d inc. direct & extr. 	100% correct ID after max. 14d inc.
	Calderaro <i>et al.</i> , 2014	<ul style="list-style-type: none"> db + extension – 24 spectra 3w inc.extr. 	correct ID after extension of db
	Karabiçak <i>et al.</i> , 2015	<ul style="list-style-type: none"> extension of db – 10 sp. 3d inc. extr. 	89.7% correct ID (<i>T. rubrum</i> complex)

MALDI-TOF MS Workflow



MALDI-TOF MS Workflow

Fast ID of the infection agent:

- Indication of the source of infection (animal or human)
- Indication of the infectivity (proper treatment)



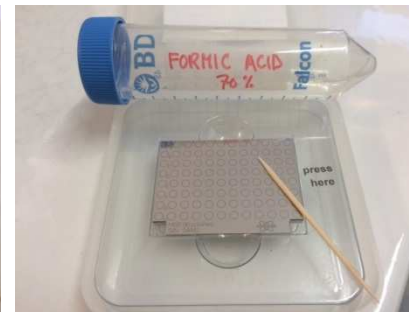
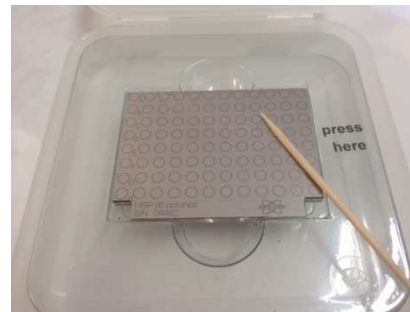
tinea capitis

(children before puberty – epidemic in school environment/family)

Tinea capitis outbreak



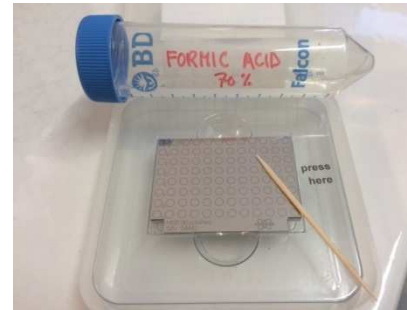
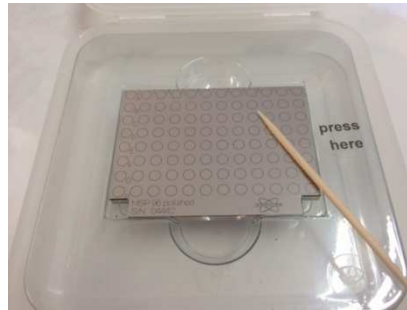
direct deposit, W and W/O 70% formic acid



Tinea capitis outbreak



direct deposit, W and W/O 70% formic acid

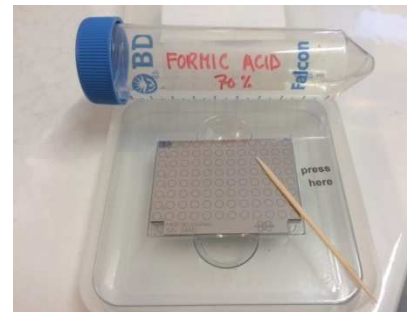
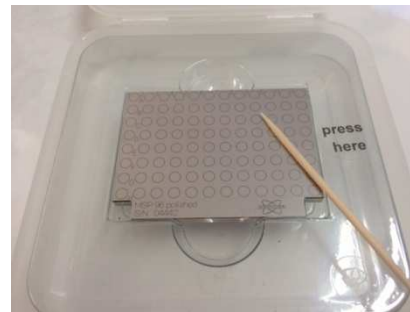


Results after first signs of growth of the dermatophytes (in validation phase)

Tinea capitis outbreak



direct deposit, W and W/O 70% formic acid



Results after first signs of growth of the dermatophytes (in validation phase)

Tinea capitis: early ID of causative agent (ID of source, initiation of a proper treatment and restriction of the outbreak)

Conclusion

PROs:

CONS:



Conclusion

PROs:

- User friendly and robust
- Fast and accurate identification

CONs:

Results dependent

- quality of the database (reference material)
- quality of the sample (growth stage dermatophyte)

Closely related species: difficulty to differentiate

Culture based method

Conclusion

PROs:

- User friendly and robust
- Fast and accurate identification

CONs:

Results dependent

- quality of the database (reference material)
- quality of the sample (growth stage dermatophyte)

Closely related species: difficulty to differentiate

Culture based method

Take into account: clinical, macro- and microscopic features

Acknowledgements



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Thank you for your attention!

