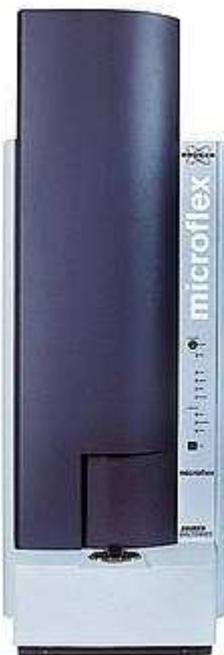


MALDI-TOF/MS

(Matrix-Assisted Laser Desorption Ionization–Time
of Flight Mass Spectrometry)

en bactériologie clinique



A. Sami CHERKAOUI
Bacteriology Lab
Service of Infectious Diseases
Geneva University Hospitals



Patiente de 25 ans

- Consulte son MT Jeudi 19.02 pour angine avec EF
- StrepA rapid-test: nég
- TTT Brufen-Dafalgan
- Samedi 21.02: CAU pour persistance angine+EF
 - Suspicion d'abcès rétro-amygdalien
 - CT-scan en urgence, hémocs puis Augmentin po

Evolution

- Lundi 23.02: hémoc positive BG-
- ID par tests biochimiques / PCR Séquençage : 24h-72h
- Même jour, *Fusobacterium necrophorum* ID par MALDI-TOF/MS, la patiente est rentrée à domicile... »
- Décision de rappeler son médecin traitant pour qu'il contacte la patiente: suspicion de Lemierre?

Patient de 67 ans

- Un patient de 67 ans est admis aux HUG suite à une altération de l'état général, 39°C, Antibio X
- Une paire d'hémoculture a été faite aux urgences plus d'autres examens
- Après 18h d'incubation un des deux flacons d'hémoculture s'est positivé
- L'examen direct montre au jour 0: Bâtonnets GRAM négatifs fusiformes

ID Phénotypique

- Jour 1 : colonies « translucide » sur gélose au sang et chocolat agar, oxidase –

- Jour 2: Résultat Vitek2:
 - Good identification
 - %id= 89 *Sphingomonas paucimobilis*

Jour 3

□ Résultat API 20 E:

- Bonne identification
- %id= 93 T= 0.62
- *Pasteurella pneumotropica*

□ Résultat API 20 NE:

- Bonne identification
- %id= 96.3 T=0.6
- *Mannheimia haemolytica* / *Pasteurella trehalosi*

Jour 6... ID Moléculaire

□ Résultat 16S-rDNA sequencing:

➤ *Capnocytophaga sputigena*



Retour au jour 1: MALDI-TOF/MS

Rank (Quality)	Matched Pattern	Score Value	NCBI Identifier
1 (+)	Capnocytophaga ochracea DSM 7271_DSM	1.847	1018
2 (-)	Capnocytophaga ochracea DSM 7272_DSM	1.585	1018

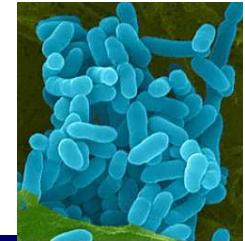
Meaning of Score Values

Range	Description
2.300 ... 3.000	highly probable species identification
2.000 ... 2.299	secure genus identification, probable species identification
1.700 ... 1.999	probable genus identification
0.000 ... 1.699	no reliable identification

Need for speed?



Need for speed?



- Monsieur H, 35 ans est admis aux HUG suite à un état fébrile, associé à des céphalées et douleur d'une loge rénale puis depuis une atteinte pulmonaire sévère avec choc septique.
- Anamnèse: Monsieur H était en vacances en Martinique, dans le village Le Vauclin.



Tempête xynthia

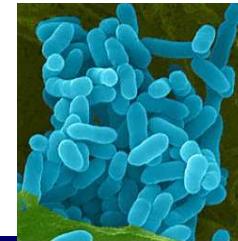
- L'état du patient se dégrade rapidement et il décède après 3 jours d'hospitalisation.

Jour 0

- Une paire d'hémoculture a été faite aux urgences plus d'autres examens

- L'examen direct montre au jour 1: Bâtonnets GRAM négatifs

Jour 1



Analyte3



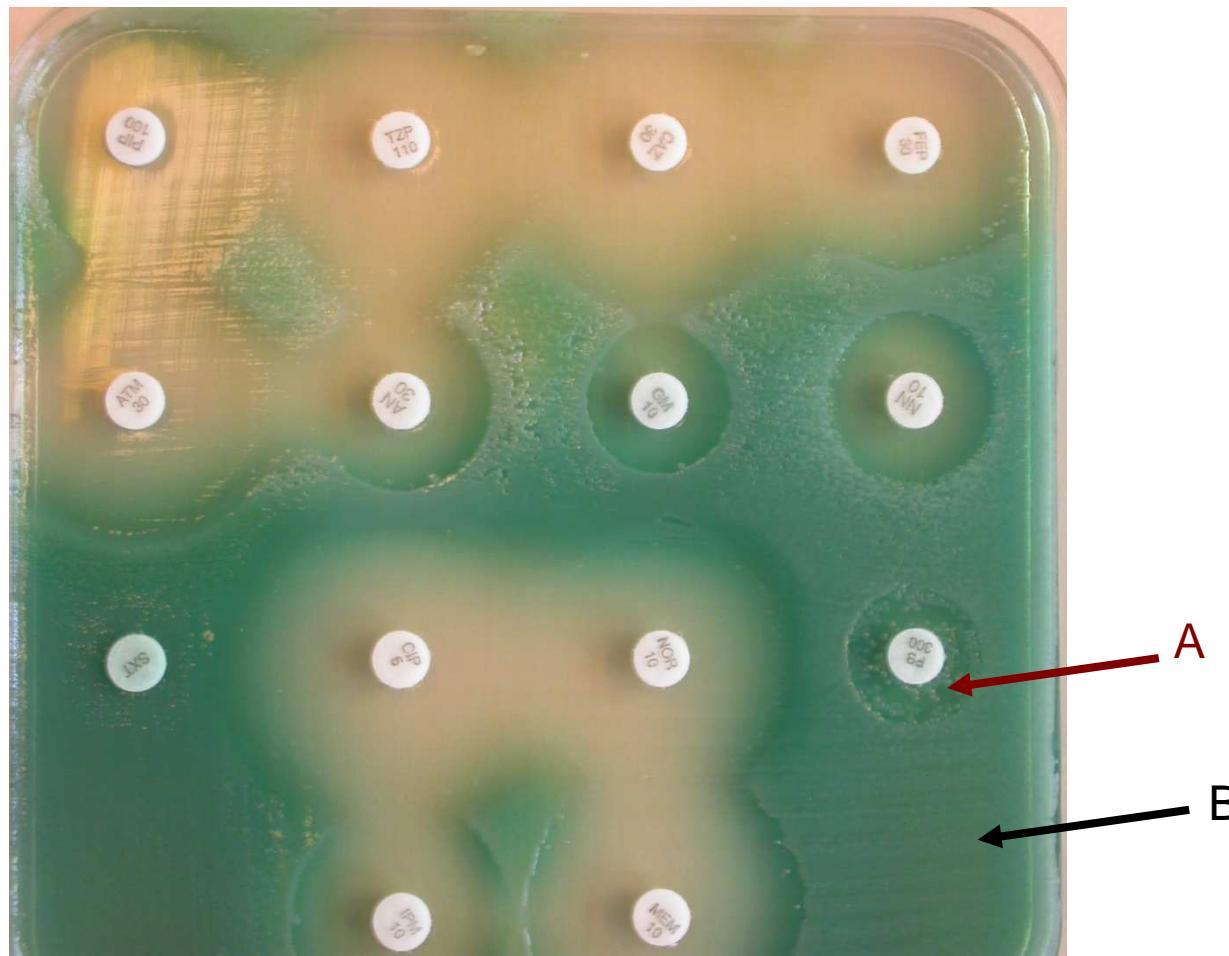
Analyte Name: 2501031407
Analyte Description:
Analyte ID: HE 9303 GS BAER
Analyte Creation Date/Time: 2010-11-26 17:17:33.375
Applied MSP Library(ies):
Applied Taxonomy Tree: Bruker Taxonomy

Rank (Quality)	Matched Pattern	Score Value	NCBI Identifier
1 (++)	Burkholderia thailandensis DSM 13276T HAM	2.045	57975
2 (-)	Burkholderia anthina LMG 16670 HAM	1.681	179879
3 (-)	Burkholderia plantarii DSM 9509T HAM	1.673	41899
4 (-)	Burkholderia cepacia group 18875_1 CHB	1.666	292
5 (-)	Burkholderia cenocepacia LMG 12614 HAM	1.655	95486
6 (-)	Burkholderia cepacia DSM 7288T HAM	1.636	292
7 (-)	Burkholderia dolosa DSM 16088T HAM	1.626	152500
8 (-)	Hydrogenophaga flava B339 UFL	1.538	65657
9 (-)	Burkholderia vietnamiensis LMG 10929T HAM	1.5	60552
10 (-)	Burkholderia ambifaria LMG 11351 HAM	1.498	152480

16S rRNA gene sequencing

- Séquence de 993pb, identique à *B. pseudomallei*.
- Une seule différence avec les 2 types de séquences de *B. mallei* connus (un C au lieu d'un T)
- Alignée avec des séquences de *B. pseudomallei*, *B. mallei* et *B. thailandensis* décrites dans l'article du JCM (Gee J.E., 2003,). La souche du patient H. présente 12 différences sur 993 bases avec *B. thailandensis*.

Antibiogramme mixte



Antibiogramme mixte

B

Identification ID:		
Date Creation Date/Time: 2008-11-20 14:51:44.39		
Used MSP Lib(s):		
Used Taxonomy Tree: Bruker Taxonomy		
Rank (Quality)	Matched Pattern	Score Value
1 (+++)	Pseudomonas aeruginosa ATCC 27853 THL	2.21
2 (++)	Pseudomonas aeruginosa 8147_2_CHB	2.18
3 (++)	Pseudomonas aeruginosa ATCC 27853_CHB	2.155
4 (++)	Pseudomonas aeruginosa 19955_1 CHB	2.062
5 (++)	Pseudomonas aeruginosa DSM 50071T HAM	2.031
6 (+)	Pseudomonas jinjuensis LMG 21316 HAM	2.003
7 (-)	Pseudomonas citronellolis DSM 50332T HAM	1.893
8 (-)	Pseudomonas indica DSM 14015T HAM	1.836
9 (-)	Lactobacillus amylovorus DSM 20552_DSM	1.799
10 (-)	Pseudomonas gessardii CIP 105469 HAM	1.72

A

Name: va 1177 dans PB		
Description:		
ID:		
Creation Date/Time: 2008-11-20 14:51:44.406		
Rank (Quality)	Matched Pattern	Score Value
1 (-)	Serratia marcescens 13103-1_CHB	2.21
2 (-)	Serratia marcescens ssp marcescens DSM 30121 DSM	2.18
3 (-)	Serratia marcescens DSM 12481 DSM	2.155
4 (-)	Serratia marcescens (PX) 24086109_MLD	2.062
5 (-)	Serratia marcescens DSM 30122 DSM	2.031
6 (-)	Serratia ureilytica DSM 16952_DSM	2.003
7 (-)	Serratia marcescens DSM 12483_DSM	1.893
8 (-)	Serratia marcescens DSM 12485 DSM	1.836
9 (-)	Serratia entomophila DSM 12358_DSM	1.799
10 (-)	Serratia marcescens ssp sakuensis CIP 107489 HAM	1.72

FinePrint - purchase at www.fineprint.com

FinePrint - purchase at www.fineprint.com

Need for speed?



Need for speed?

- Homme de 42 ans est admis au HUG suite à un état fébrile.
- Une paire d'hémoculture a été faite aux urgences plus d'autres examens
- L'examen direct montre au jour 1: CG+ en diplo et chainettes

Need for speed?

Analyte5



Analyte Name: 3001721487
Analyte Description: he 5569
Analyte ID: 2011-02-01 10:31:54.796
Analyte Creation Date/Time:
Applied MSP Library(ies):
Applied Taxonomy Tree: Bruker Taxonomy

Rank (Quality)	Matched Pattern	Score Value	NCBI Identifier
1 (++)	Streptococcus pneumoniae ATCC 49619 THL	2.04	1313
2 (++)	Streptococcus pneumoniae besSt29 THL	2.001	1313
3 (+)	Streptococcus oralis DSM 20379 DSM	1.854	1303
4 (+)	Streptococcus pneumoniae DSM 14378 DSM	1.839	1313
5 (+)	Streptococcus pneumoniae DSM 11868 DSM	1.81	1313
6 (+)	Streptococcus oralis DSM 20395 DSM	1.77	1303
7 (+)	Streptococcus pneumoniae DSM 11866 DSM	1.749	1313
8 (+)	Streptococcus pseudopneumoniae DSM 18670T DSM	1.738	257758
9 (+)	Streptococcus oralis DSM 20627T DSM	1.707	1303
10 (-)	Streptococcus pneumoniae DSM 11865 DSM	1.673	1313

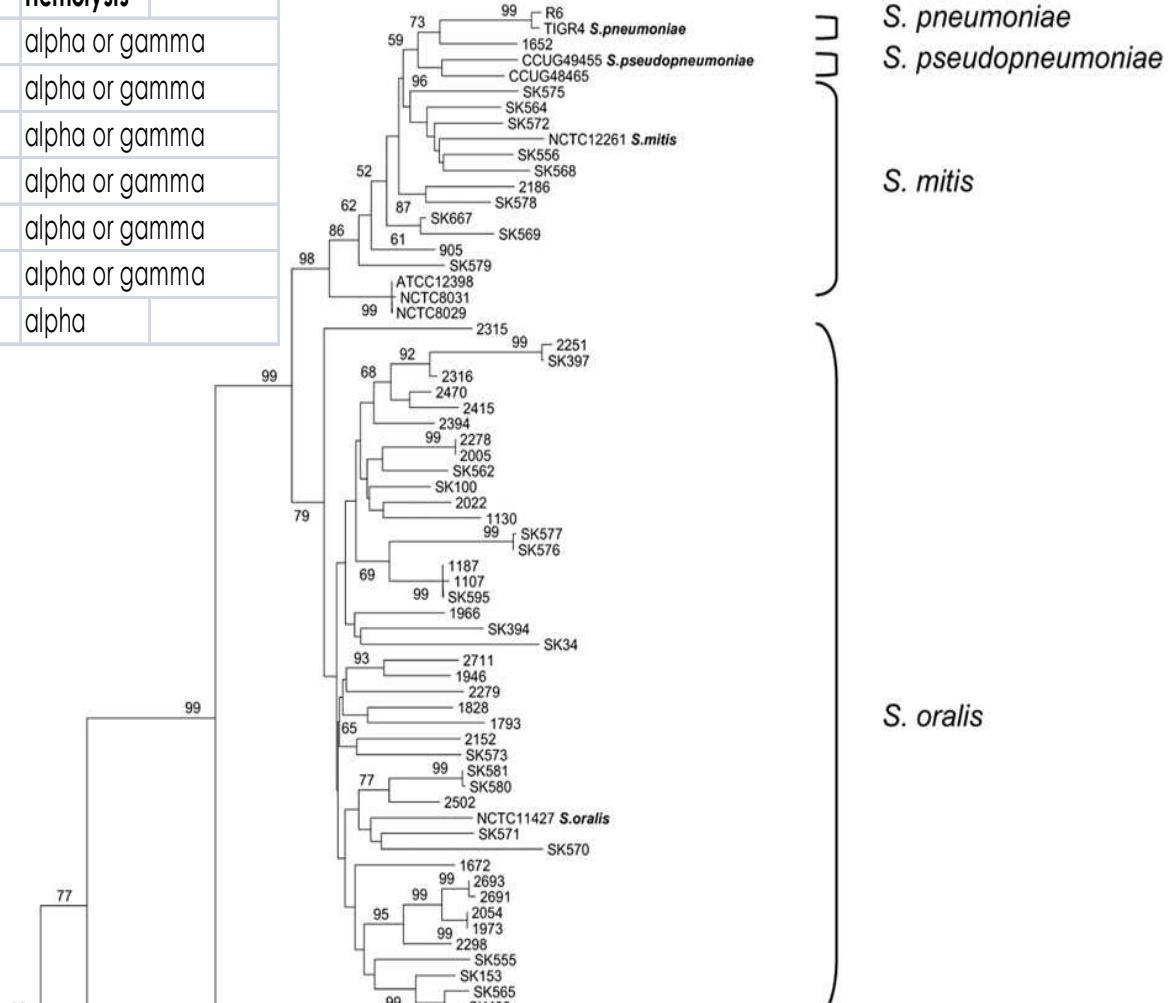
Jour 2
Optochine: Sensible
Pneumoagglutination: NEG

ID: *S. mitis*

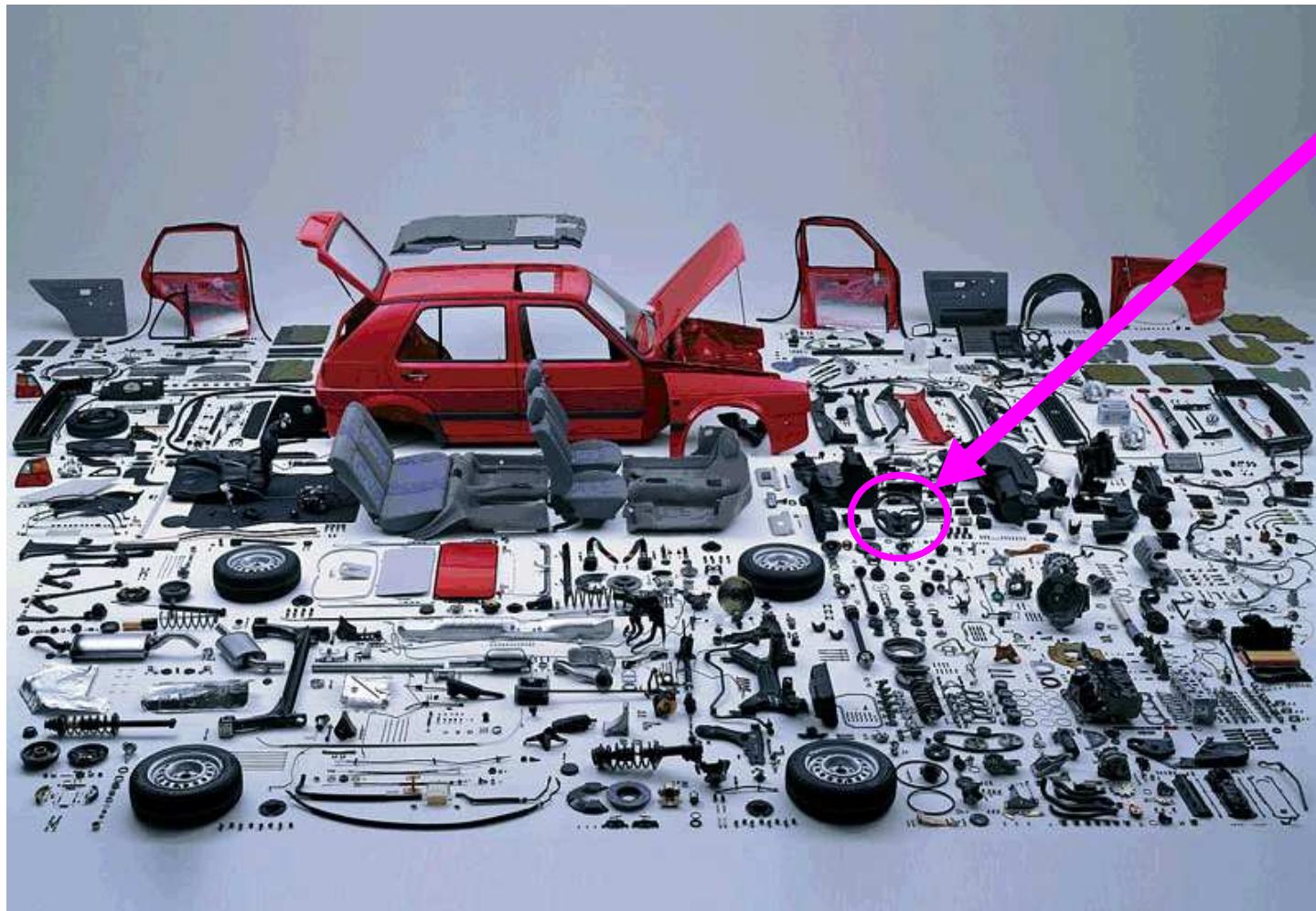
Endocardite sur
valve native

Streptococcus mitis group

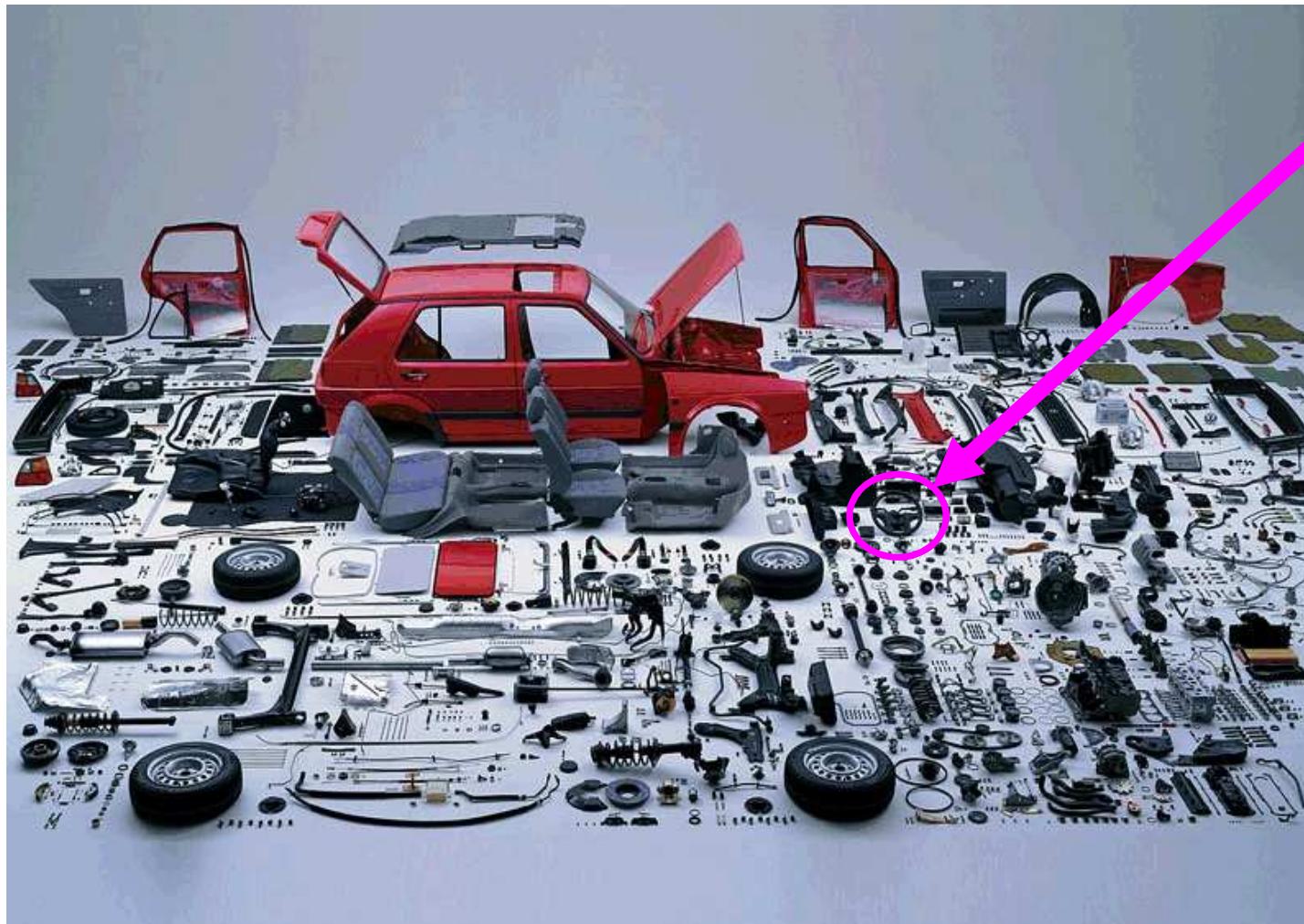
<i>Streptococcus mitis</i> group		Lancefield group	Hemolysis
<i>S. mitis</i>		K, O or None at all	alpha or gamma
<i>S. sanguinis</i>		H or None at all	alpha or gamma
<i>S. parasanguinis</i>		None at all	alpha or gamma
<i>S. oralis</i>		None at all	alpha or gamma
<i>S. infantis</i>		None at all	alpha or gamma
<i>S. gordonii</i>		None at all	alpha or gamma
<i>S. pneumoniae</i>		None at all	alpha



The principle of MS...

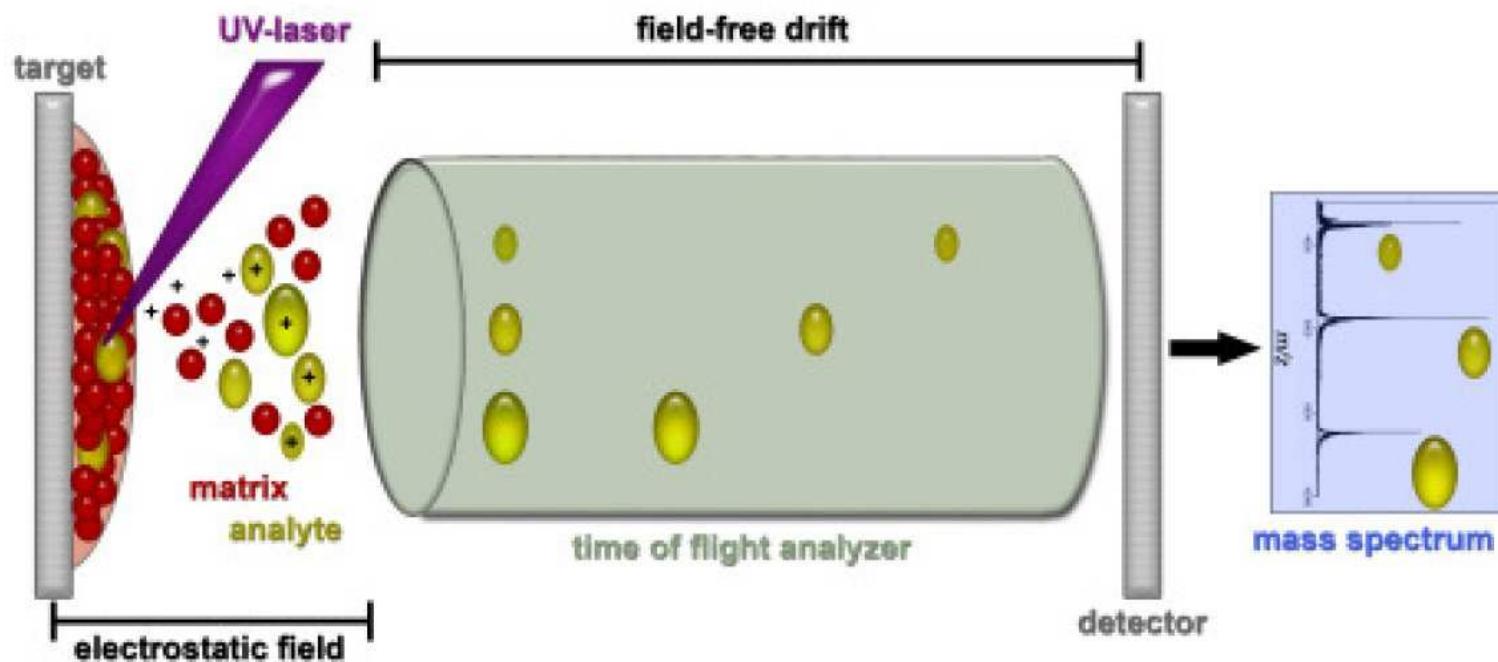


The principle of MS...

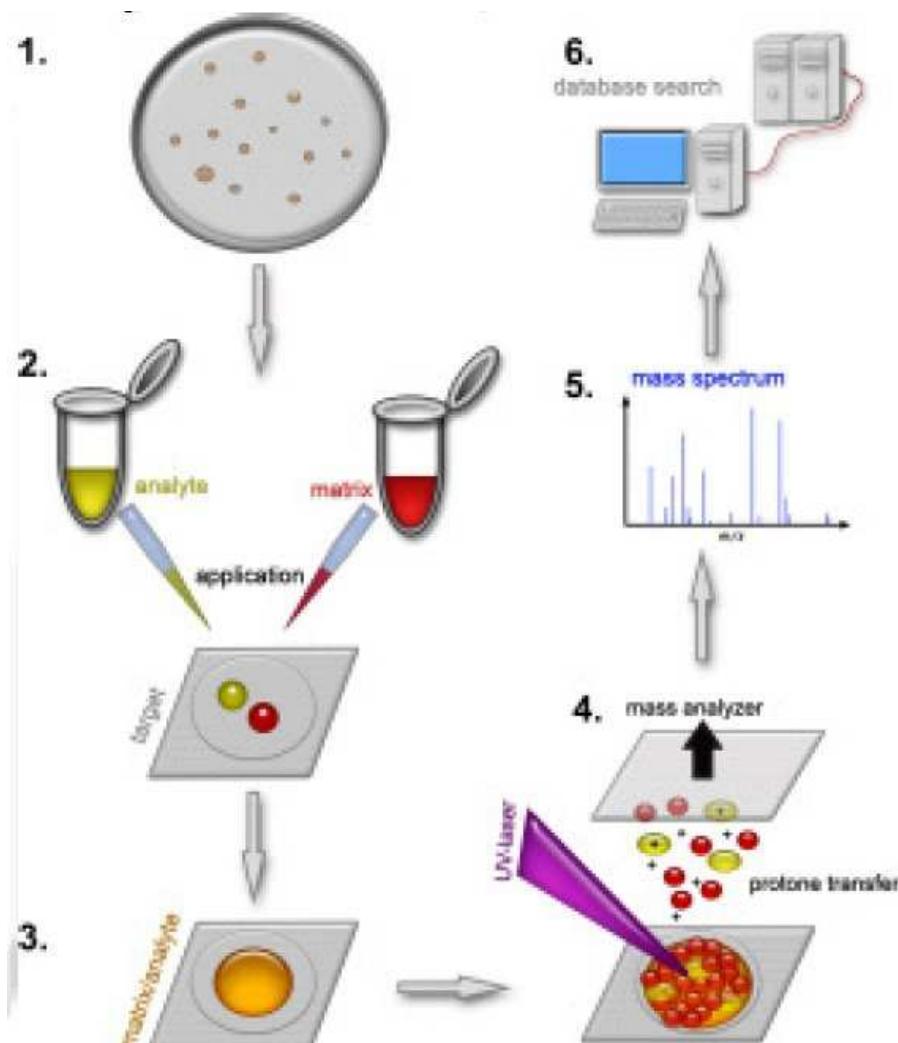


This was
a car!

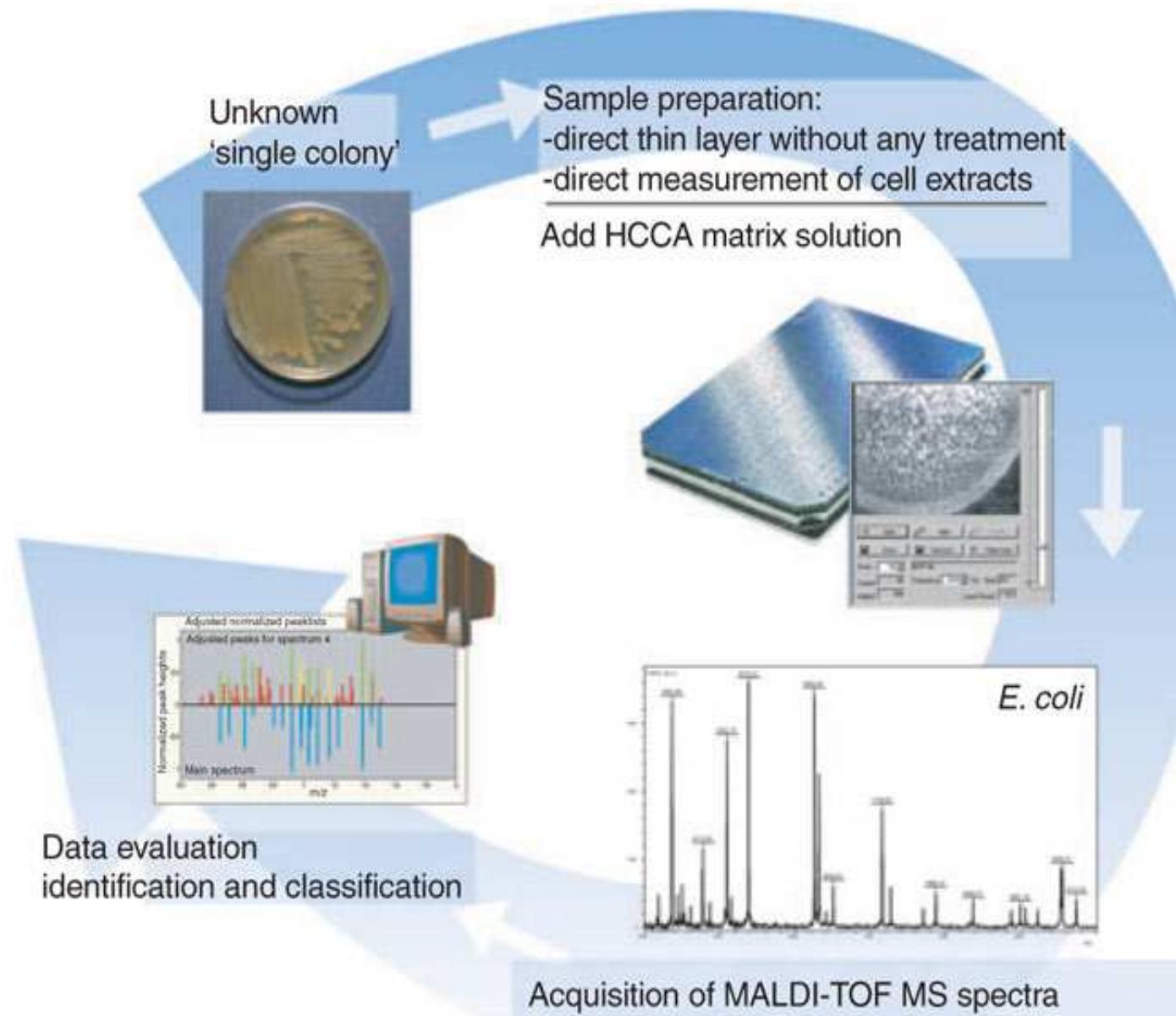
The principle of MS...



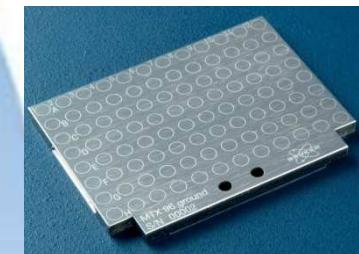
Workflow of MALDI-TOF MS-based bacterial identification



α -cyano-4-hydroxy-cinnamic acid in 50% acetonitrile-
2.5% trifluoroacetic acid

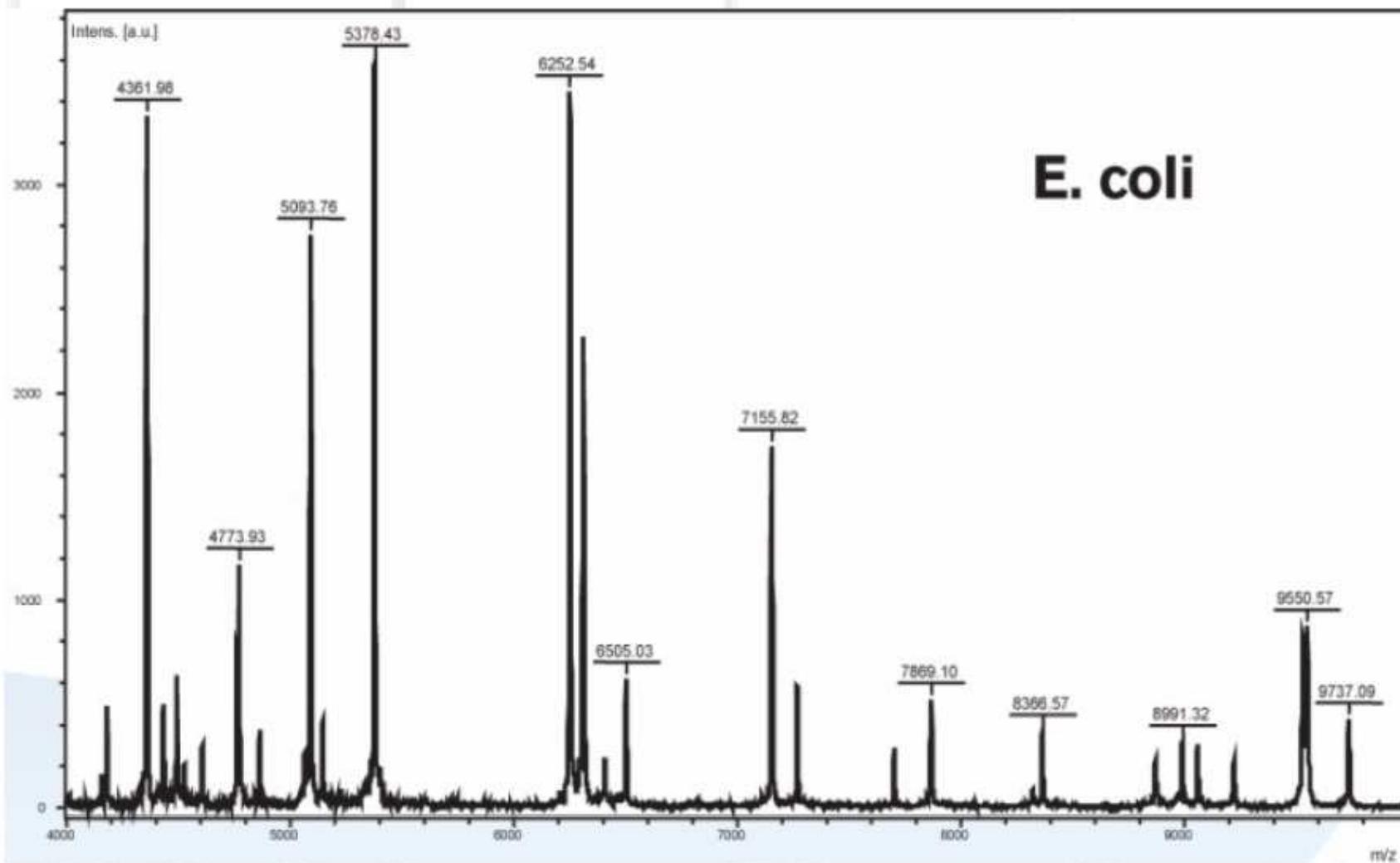


Put one colony
on the target
And add a matrix

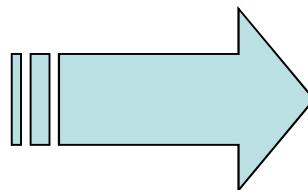
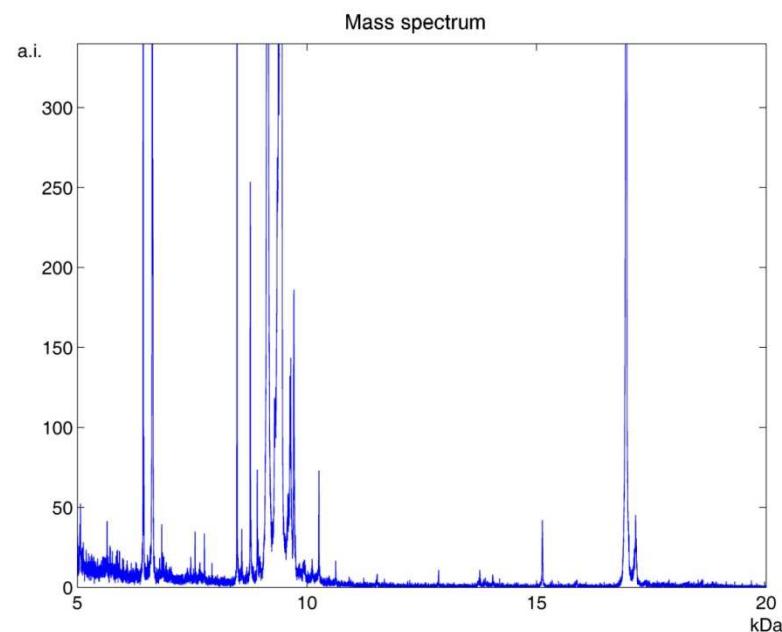


Insert the target
and start shooting

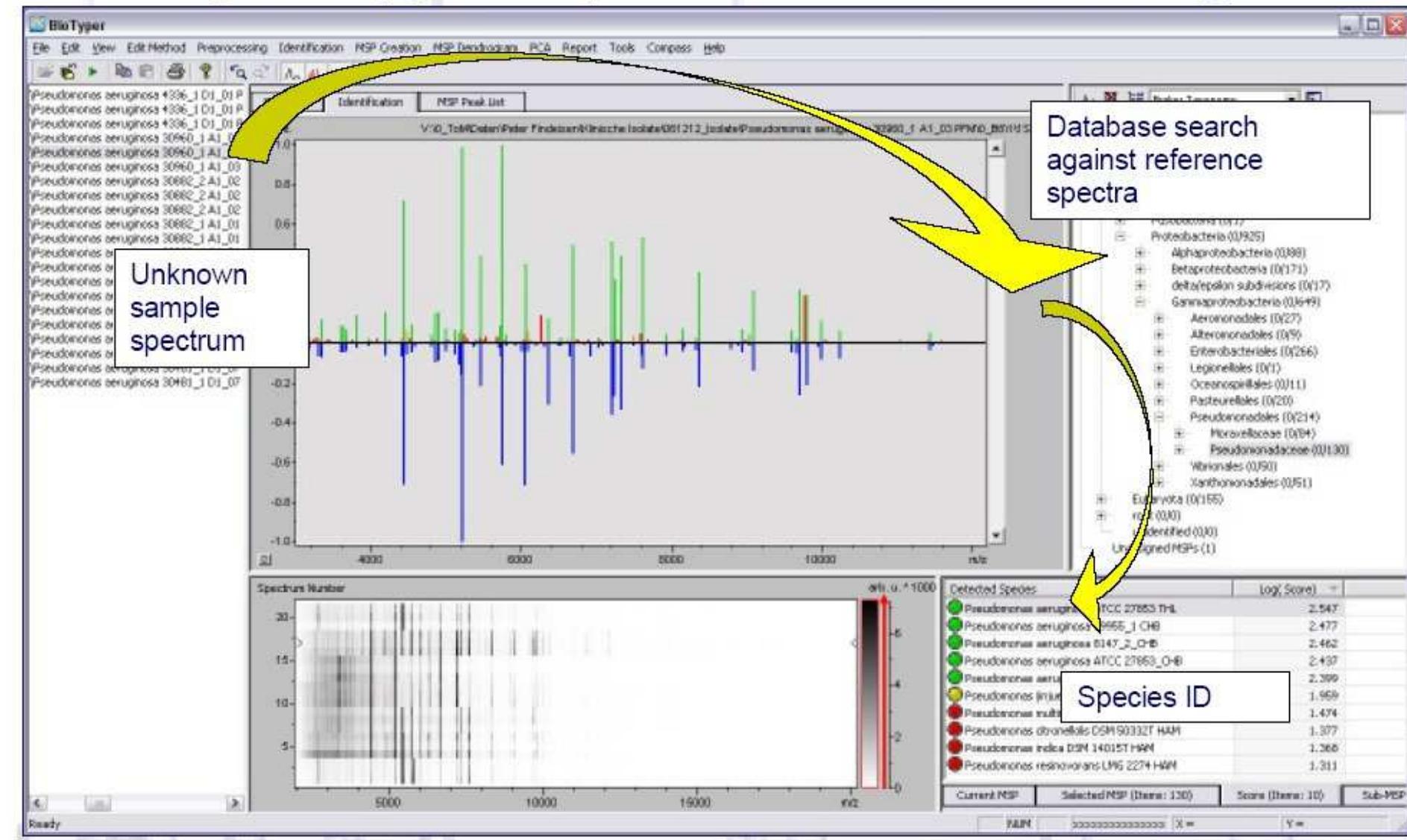
Example of a MALDI-TOF MS Spectrum



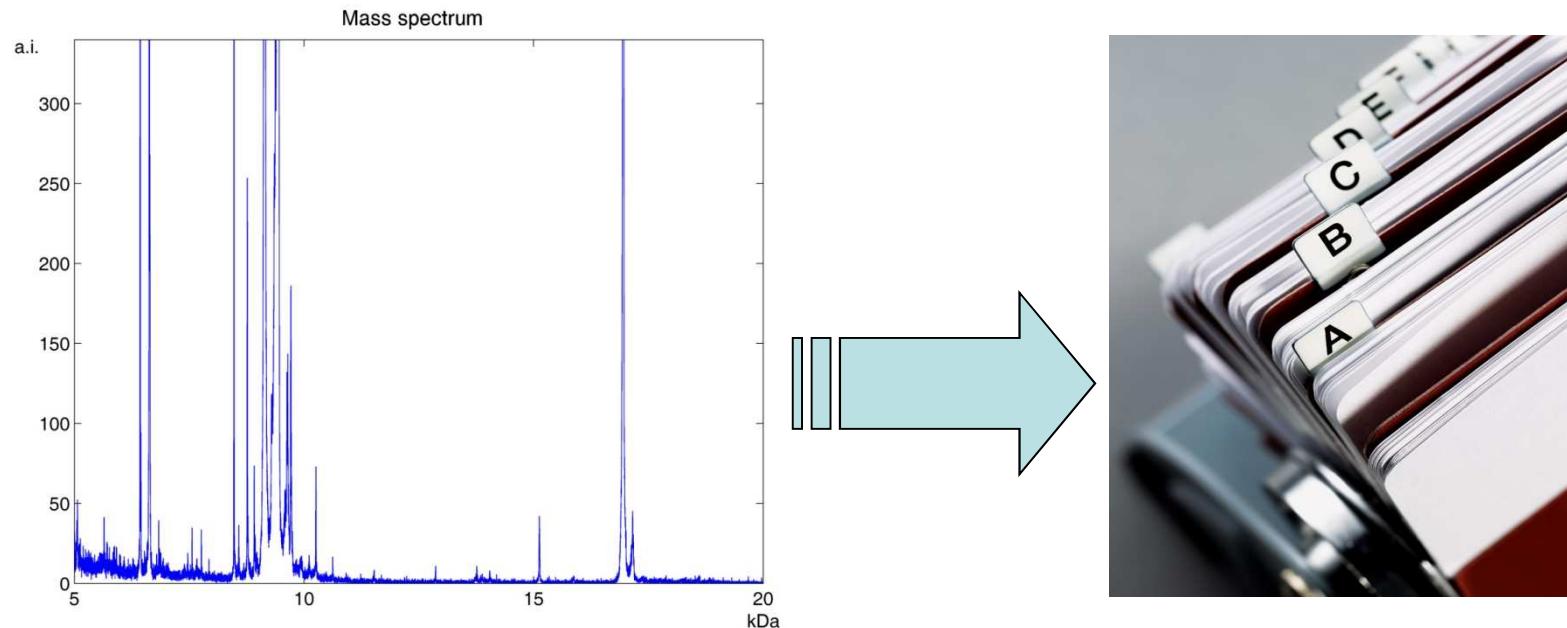
Data are then interpreted



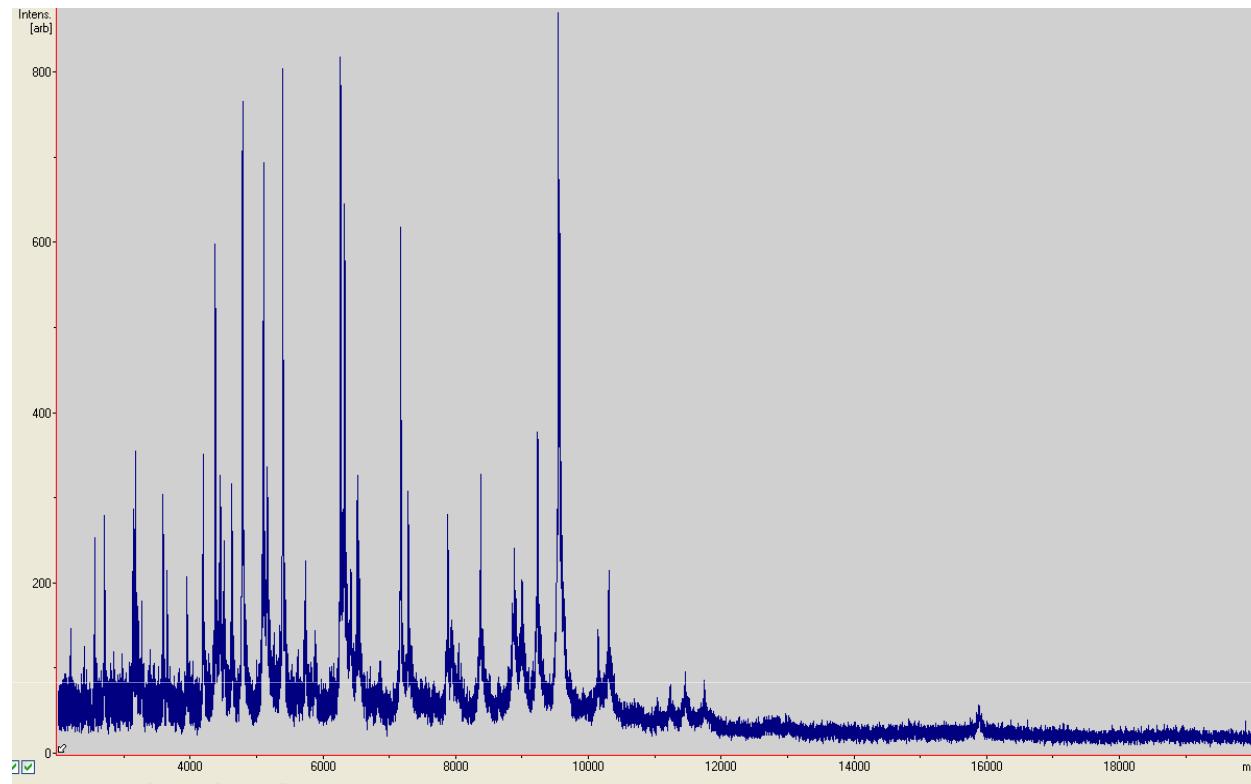
MALDI-TOF MS Analysis Software (BioTyper™) – Pattern Matching



Data are then interpreted



Range	Description	Symbols	Color
2.300 ... 3.000	highly probable species identification	(+++)	green
2.000 ... 2.299	secure genus identification, probable species identification	(++)	green
1.700 ... 1.999	probable genus identification	(+)	yellow
0.000 ... 1.699	no reliable identification	(-)	red



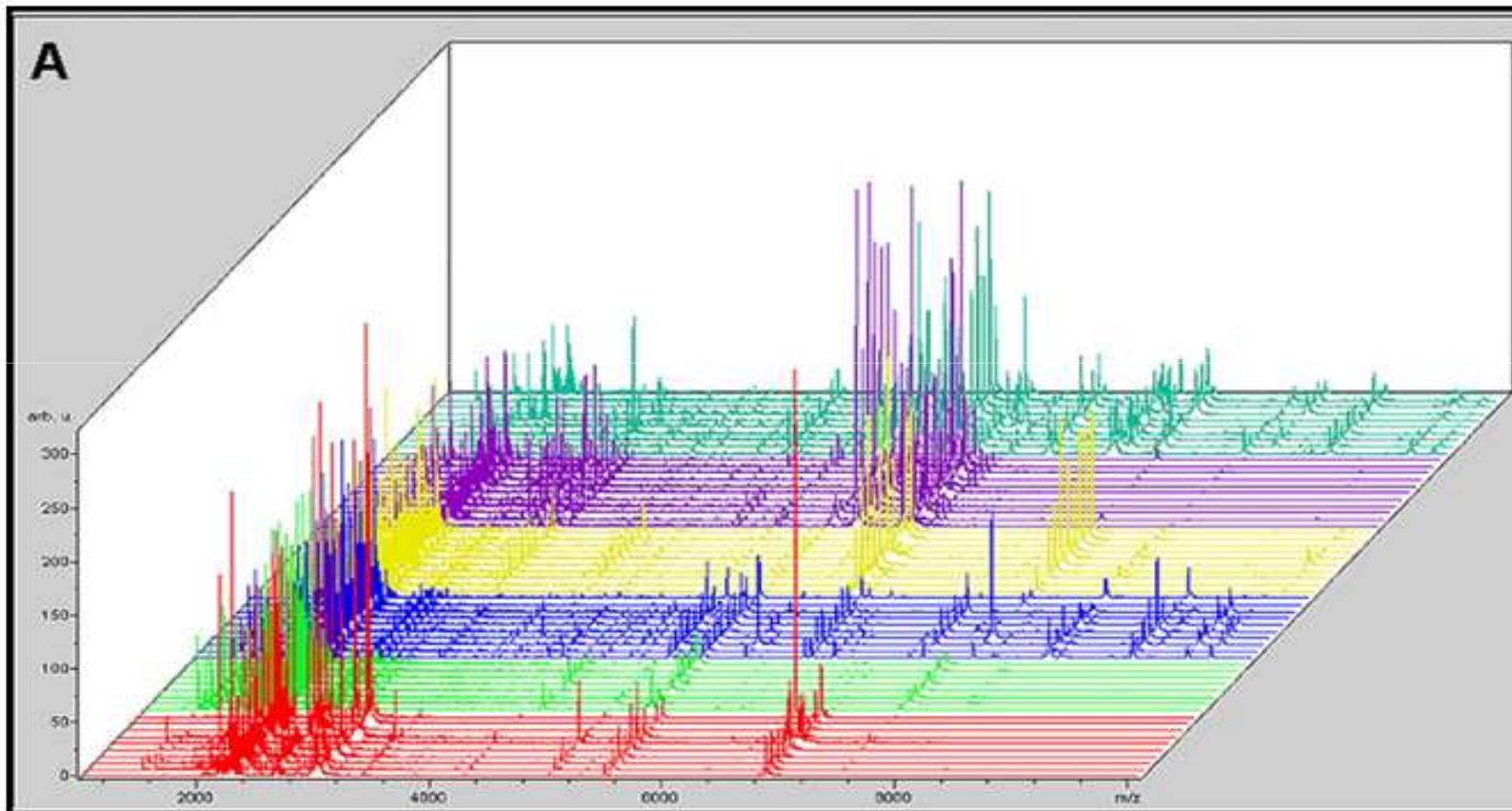
Result Overview

Analyte Name	Analyte ID	Organism (best match)	Score Value	Organism (second best match)	Score Value
ECOL ATCC 25922 (+++)		Escherichia coli	2.474	Escherichia coli	2.376

Meaning of Score Values

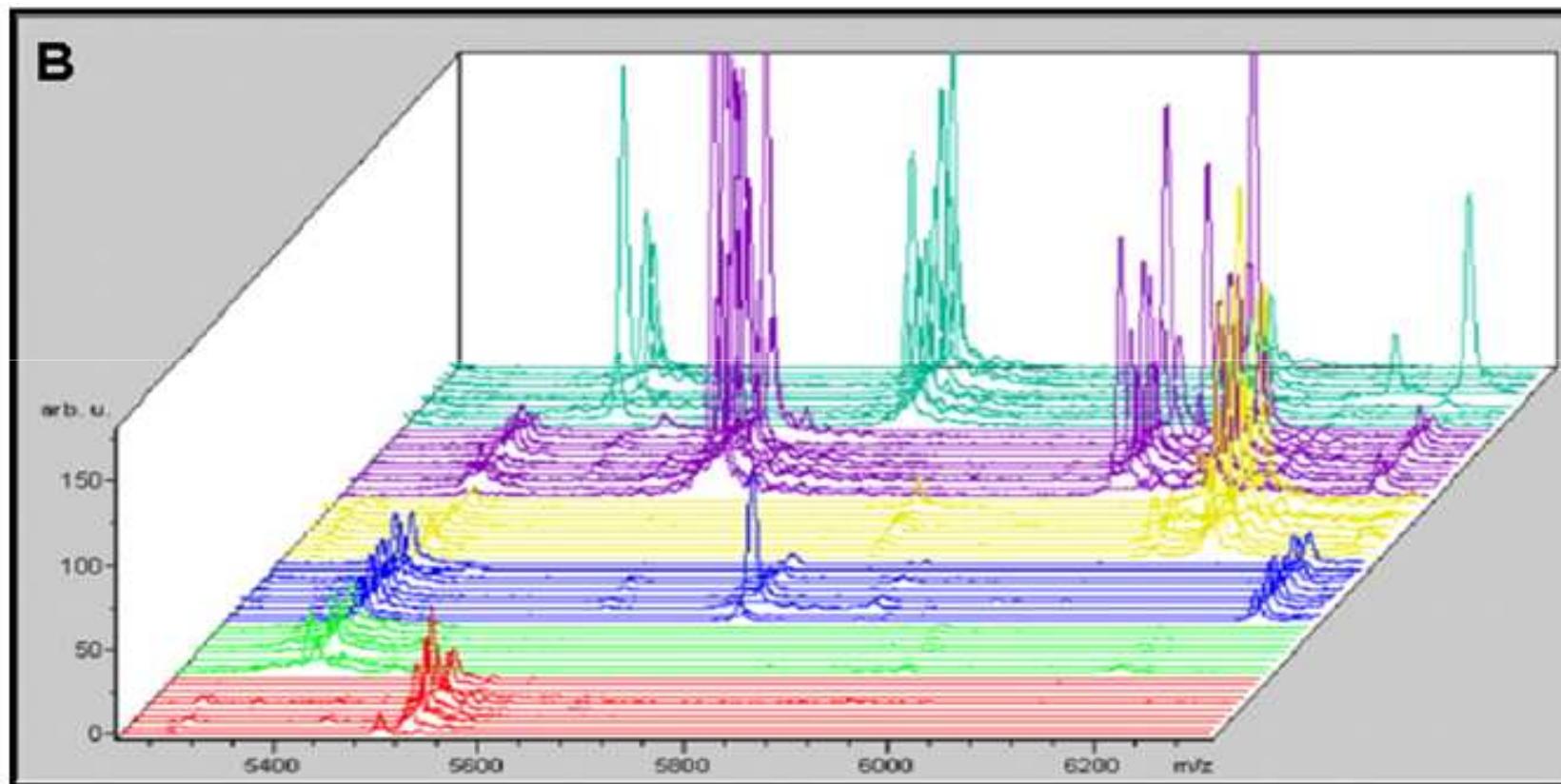
Range	Description	Symbols	Color
2.300 ... 3.000	highly probable species identification	(+++)	green
2.000 ... 2.299	secure genus identification, probable species identification	(++)	green
1.700 ... 1.999	probable genus identification	(+)	yellow
0.000 ... 1.699	not reliable identification	(-)	red

The principle of MS...



S. aureus; *Streptococcus* group B; *E. coli*; *K. pneumoniae*; *Salmonella* serotype B; and, *P. aeruginosa* over m/z 1800–9000

The spectra are enlarged in the region of 5000–8500 Da/ z



S. aureus; *Streptococcus* group B; *E. coli*; *K. pneumoniae*; *Salmonella* serotype B; and, *P. aeruginosa* over m/z 1800–9000

Comparison of Two Matrix-Assisted Laser Desorption Ionization–Time of Flight Mass Spectrometry Methods with Conventional Phenotypic Identification for Routine Identification of Bacteria to the Species Level^V

Abdessalam Cherkaoui,^{1,*} Jonathan Hibbs,² Stéphane Emonet,¹ Manuela Tangomo,² Myriam Girard,²
Patrice Francois,² and Jacques Schrenzel^{1,2}

*Clinical Microbiology Laboratory¹ and Genomic Research Laboratory², Service of Infectious Diseases,
University of Geneva Hospitals (HUG), CH-1211 Geneva 14, Switzerland*

Received 23 September 2009/Returned for modification 23 November 2009/Accepted 9 February 2010

Study design

- 720 clinical isolates
- Collected during 21 consecutive working days
- Various swabs (n=261), urine (n=197), blood cultures (n=120), respiratory tract (n=120), and stools (n=12)

Study design

- 720 clinical isolates
- Collected during 21 consecutive working days
- Various swabs (n=261), urine (n=197), blood cultures (n=120), respiratory tract (n=120), and stools (n=12)
- Two rotating laboratory technologists processed in parallel freshly isolated colonies on two MS systems
- Results were first compared to our phenotypic routine (API and Vitek2)
- Any discordant result led to 16S rDNA sequencing (gold standard)

Interpretation of the results

- 1- high score value ≥ 2.0 for Bruker and ≥ 90 for Shimadzu
=> isolates rated as identified to the species level

- 2- intermediate score value ≥ 1.7 for B and ≥ 70 for S
=> isolates rated as identified to the genus level

- 3- low score value < 1.7 for B and < 70 for S
=> isolates rated as ambiguous, thus not reported

Study design (2)

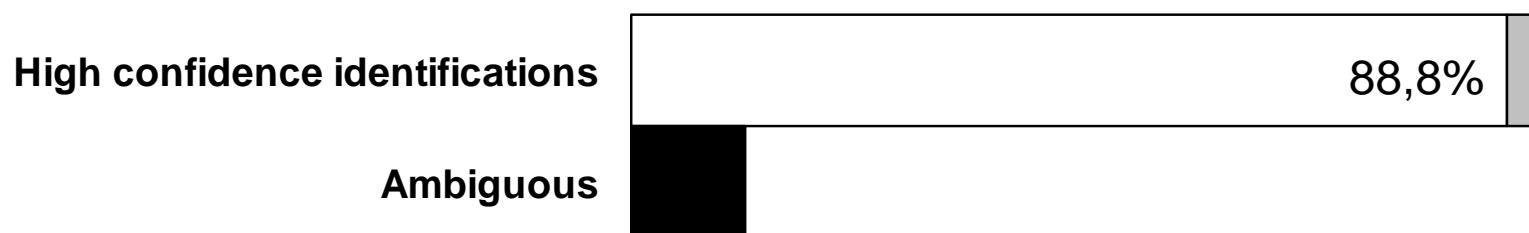
- Whenever the quality of MS scores revealed ambiguous, a second experiment was performed on a fresh bacterial culture.
 - Additional identification was performed on 54/720 (7.5%) and 39/720 (5.4 %) isolates, on MS 1 and 2

- No attempt was made to employ the specific extraction procedures recommended by the manufacturers, e.g. for streptococcal speciation.

Accuracy of MALDI-TOF MS identifications of 720 clinical isolates

Shimadzu

- Concordant with conventional methods**
- Concordant with PCR**
- Incorrect or ambiguous**

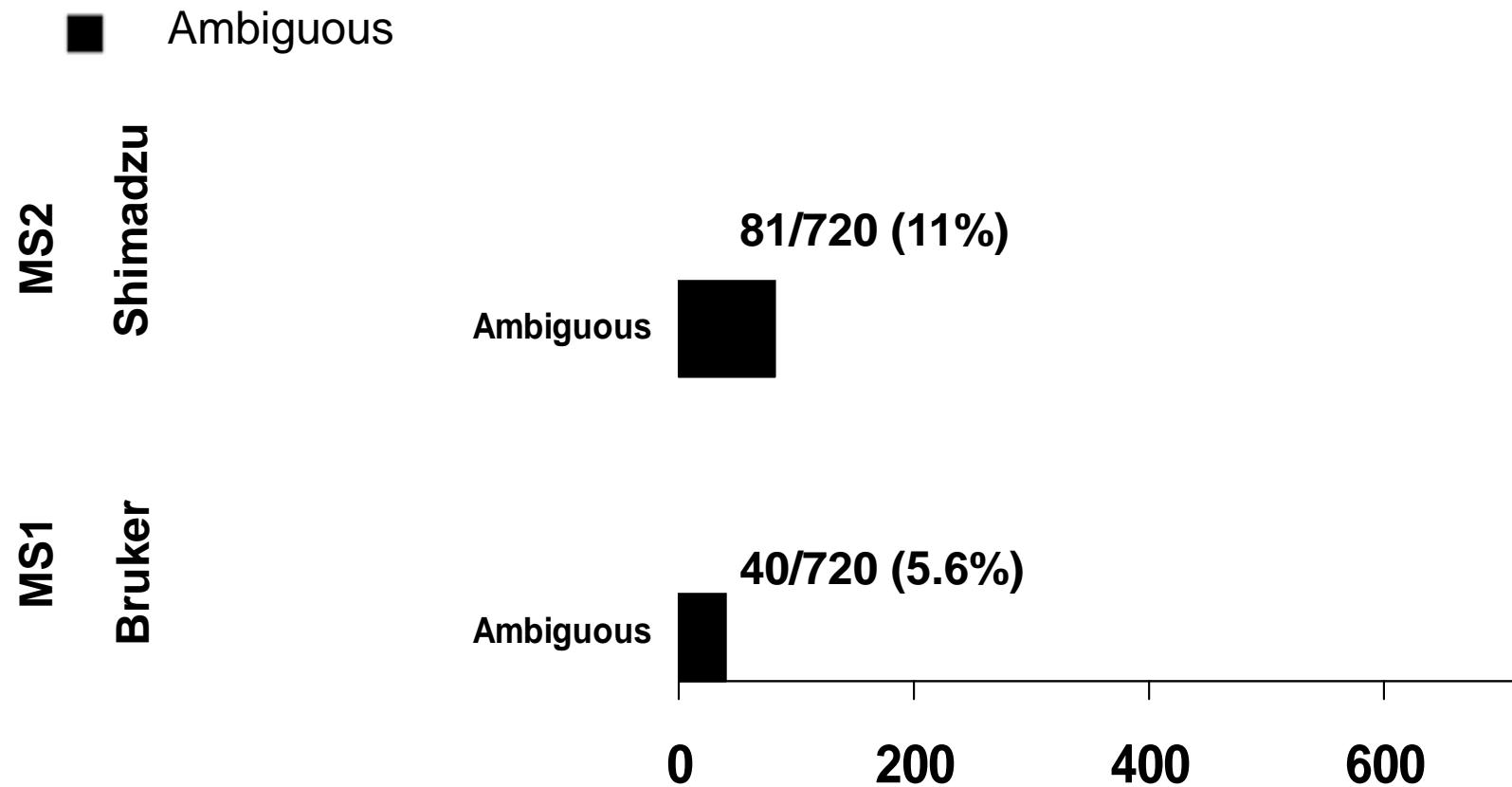


Bruker



0 200 400 600

Accuracy of MALDI-TOF MS id of 720 clinical isolates



Analysis of all incorrect cases

Cases	Phenotypic	16S rDNA	MALDI-MS	System
1	No ID	Achromobacter xylosoxidans	A. denitrificans	Bruker
1	S. epid	No ID	S. capitis	Bruker
1	S. mitis	Streptococcus sp	S. pneumoniae	Bruker
2	No ID	Streptococcus bovis	S. lutiensis	Bruker
1	No ID	Clostridium boltei	C. casei	Bruker
1	S. epid	Staphylococcus sp	S. capitis/caprae	Shimadzu
1	S. mitis	Streptococcus sp	S. pneumoniae	Shimadzu
1	S. mitis	Streptococcus sp	S. oralis	Shimadzu

Analysis of all incorrect cases

Cases	Phenotypic	16S rDNA	MALDI-MS	System
1	No ID	Achromobacter xylosoxidans	A. denitrificans	Bruker
1	S. epid	No ID	S. car	Bruker
1	S. mitis	Streptococcus sp		Bruker
2	No ID	Streptococcus bo		Bruker
1	No ID	Clostridium boltei		Bruker
1	S. epid	Staphylococcus sp		Shimadzu
1	S. mitis	Streptococcus sp	S. pneumoniae	Shimadzu
1	S. mitis	Streptococcus sp	S. oralis	Shimadzu

Establish
rules
considering
these cases

Results: based on species

- Outstanding identification of Enterobacteriaceae,
Staphylococcus aureus and *Pseudomonas aeruginosa*.
- Lower yields (but no false-positive identification) when analyzing enterococci (75-56%) or streptococci (37-26%) to the species level.
- Latter results could certainly be improved by using dedicated matrix.

Results: four categories

- 1- correct identification (same as biochemical tests) with a high score value ≥ 2.0 for Bruker and ≥ 90 for Shimadzu;
=> isolates rated as identified to the species level
- 2- correct identification (same as biochemical tests) with an intermediate score value ≥ 1.7 for B and ≥ 70 for S;
=> isolates rated as identified to the genus level
- 3- MS results inferior to the biochemical identification
- 4- MS results superior to the biochemical identification

Results: four categories

1- correct identification - identified to the species level:
620/720 (86%) and 559/720 isolates (78%)

2- correct identification - identified to the genus level:
46/720 (6%) and 95/720 isolates (13%)

3- MS results inferior to the biochemical identification:
18/49 (37%) and 28/49 (57%), **but none wrong!**

4- MS results superior to the biochemical identification:
15/49 (30%) and 11/49 (23%) to the species level
16/49 (33%) and 10/49 (20%) to the genus level

Cost and timeliness estimates of conventional identification

	<u>Cost per isolate (\$US)</u>		<u>Turnaround time (hr)</u>	
	Avg.	Total	Avg.	Total
<i>E. coli</i> (n=216)	\$0.20	\$43	1	216
<i>S. aureus</i> (n=55)	\$1.50	\$83	1	55
Other (n=449)	\$10.00	\$4,490	24	10,776
All isolates (n=720)	Total cost:	\$4,616	Average time: 15 hrs	

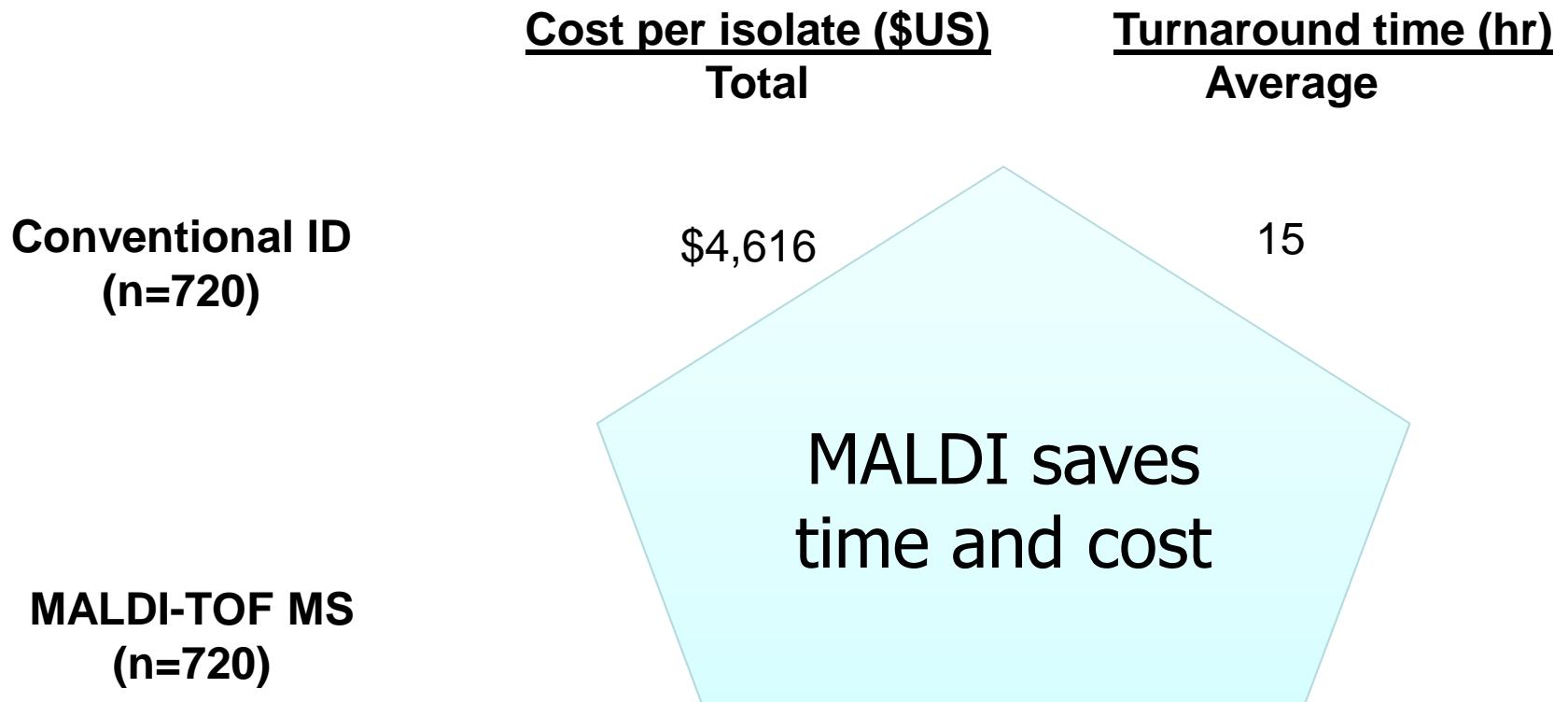
Cost and timeliness estimates of Bruker-based identification

	<u>Cost per isolate (\$US)</u>		<u>Turnaround time (hr)</u>	
	Avg.	Total	Avg.	Total
High confidence MALDI-TOF MS (n=636)	\$0.50	\$318	0.08	53
Lower-confidence and Ambiguous MALDI-TOF MS (n=84)	\$10.50	\$882	24	2,016
Total cost:		\$1,200	Average time: 3 hrs	

Cost and timeliness estimates of identification methods

	<u>Cost per isolate (\$US)</u>	<u>Turnaround time (hr)</u>
	Total	Average
Conventional ID (n=720)	\$4,616	15
MALDI-TOF MS (n=720)	\$1,200	3

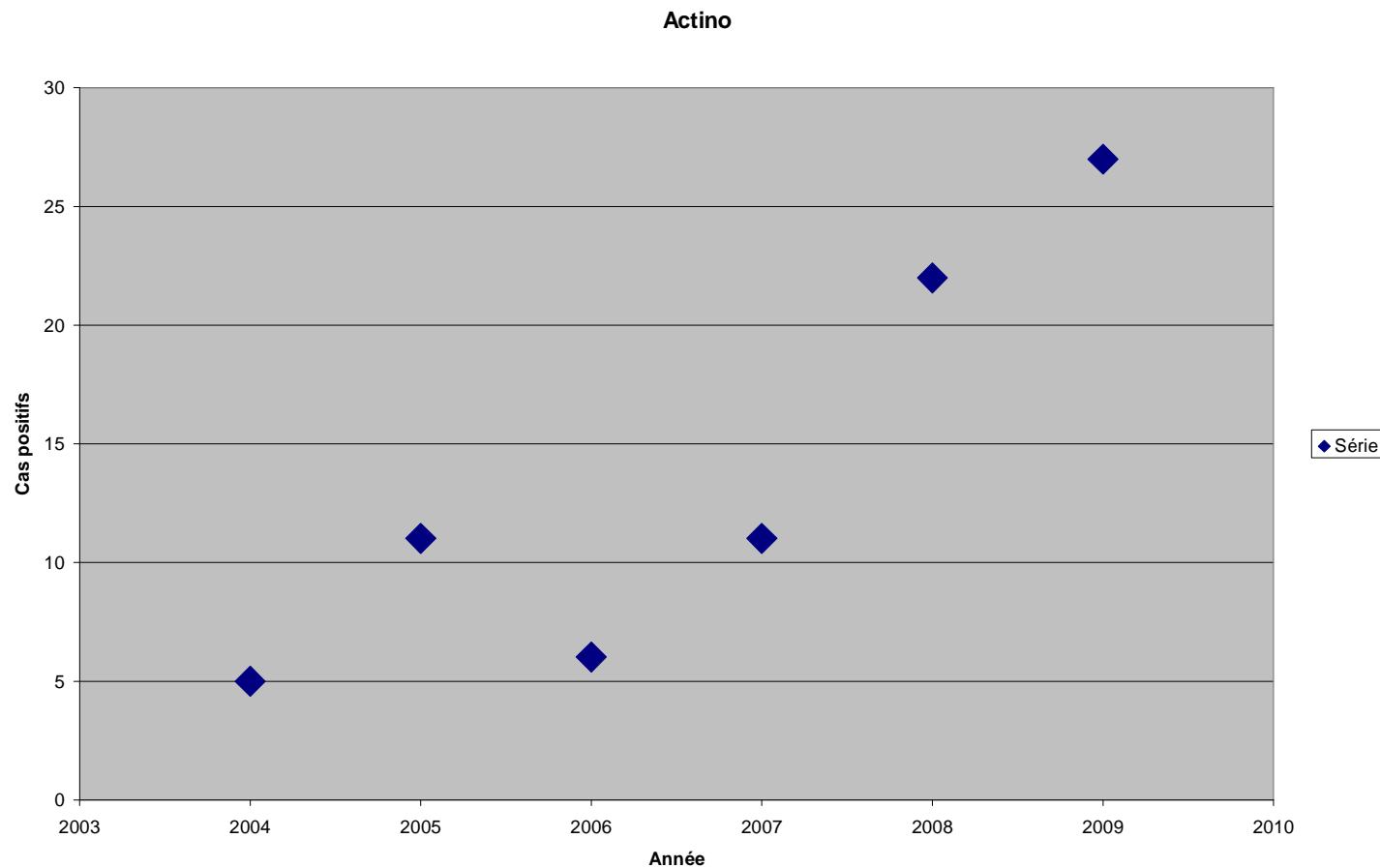
Cost and timeliness estimates of identification methods



Evaluation of matrix-assisted laser desorption/ionization time-of-flight mass spectrometry for the rapid identification of beta-hemolytic *streptococci*

- MALDI-TOF/MS analysis of 386 sub-cultured beta-hemolytic *streptococci* isolates resulted in high-confidence identifications to the species level for all 386 isolates, with 100% concordance with the Lancefield group classification as determined by the latex agglutination kit.
- The Vitek2 gave high-confidence identification to the species level for 269/306 isolates of group B streptococci (88%), 48/52 isolates of group A streptococci (92%) and 11/28 isolates of group C and G streptococci (39%).
- Fourteen isolates were not identified by Vitek2, 12 were incorrectly identified and 32 isolates were identified ambiguously (%ID=50) as *Streptococcus agalactiae* or *Streptococcus dysgalactiae*.
- Whenever high-confidence MS identifications disagreed with Vitek2 results, 16S sequencing resolved identifications in favor of the MS system.
- The 12 incorrect Vitek2 results (%ID>90) were as follows: six isolates identified as *Gemella haemolysans*, two isolates identified as *Kocuria rosea*, one isolate identified as *Lactobacillus gravieae*, one isolate identified as *Streptococcus pluranimalium* and one isolate identified as *Kocuria varians*.

Actinomyces spp. Identifiés au LCB: l'effet « MALDI-TOF/MS »...



Implementation: a few steps

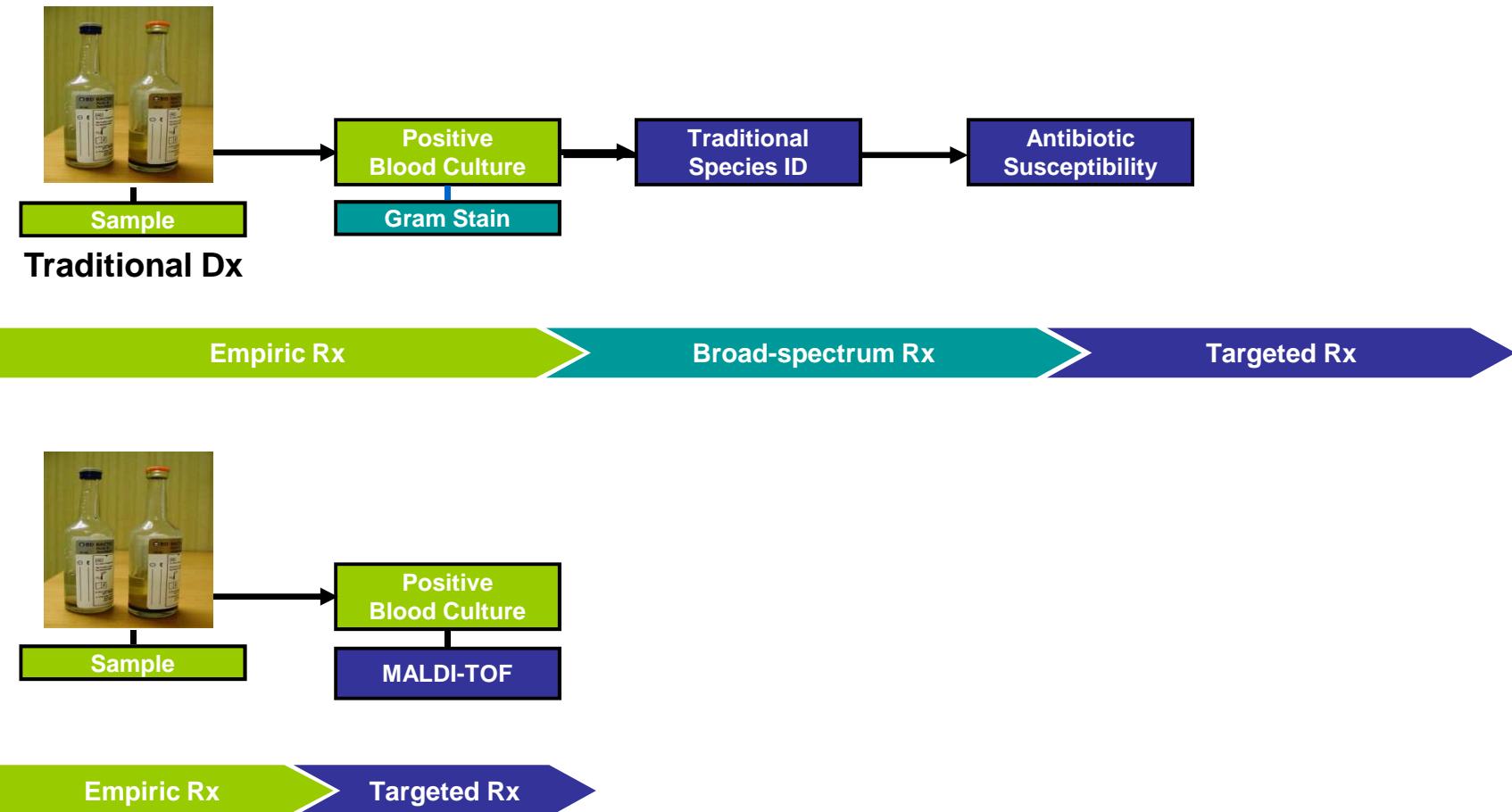
Organismes	Gram/autres	Tests
<i>S. aureus</i>	Cg+ amas	Catalase, Pastorex, DNase, Coagulase <i>mec A et femA,</i>
<i>S. pneumoniae</i>	Cg+ diplocuques	Optochine , Agglut. Pneumokit
<i>S. pyogenes</i> <i>S. agalactiae</i> <i>S. dysgalactiae</i>	Cg+ chaînes	Bacitracine , Bactrim Slidex Streptokit
Enterococcus spp.	Cg+ chaînes	Bile esculine, NaCl 6.5%, Tellurite Galerie AP / Vitek2
<i>H. influenzae</i>	Petits bg-	Porphyrine , Galerie AP
<i>M. catarrhalis</i>	Coccobacilles Gram -	Oxidase,Tributyrine
Entérobactéries	Fermentation du Lactose, Oxidase	Vitek2 / Galerie AP
Germes non fermentatifs	Fermentation du Lactose, Oxidase	Vitek2 / Galerie AP
<i>B. cepacia</i>	Fermentation du Lactose	Oxidase variable Vitek2 / Galerie AP

Implementation: a few steps

Organismes	Gram/autres	Tests
<i>S. aureus</i>	Cg+ amas	Catalase, Pastorex, DNase, Vitek, Vitek2, mecA
<i>S. pneumoniae</i>	Cg+ diplocuques	Optochin, Vitek, Vitek2, Pneumococcal agglut.
<i>S. pyogenes</i> <i>S. agalactiae</i> <i>S. dysgalactiae</i>	Cg+ chaînes	Bacitracine, Vitek, Vitek2, Bacitracine, Vitek, Vitek2, Slide, Vitek, Vitek2
Enterococcus spp.	Cg+ chaînes	Bile escharique, Vitek, Vitek2, 6.5% NaCl, Vitek, Vitek2, Galei, Vitek, Vitek2
<i>H. influenzae</i>	Petits bg-	Porphyrines, Vitek, Vitek2, AP
<i>M. catarrhalis</i>	Coccobacilles Gram -	Oxidase, Vitek, Vitek2, AP
Entérobactéries	Fermentation du Lactose, Oxidase	Vitek, Vitek2, Vitek, Vitek2, AP
Germes non fermentatifs	Fermentation du Lactose, Oxidase	Vitek, Vitek2, Vitek, Vitek2, AP
<i>B. cepacia</i>	Fermentation du Lactose	Oxidase, Vitek, Vitek2, Vitek, Vitek2 / Galerie AP

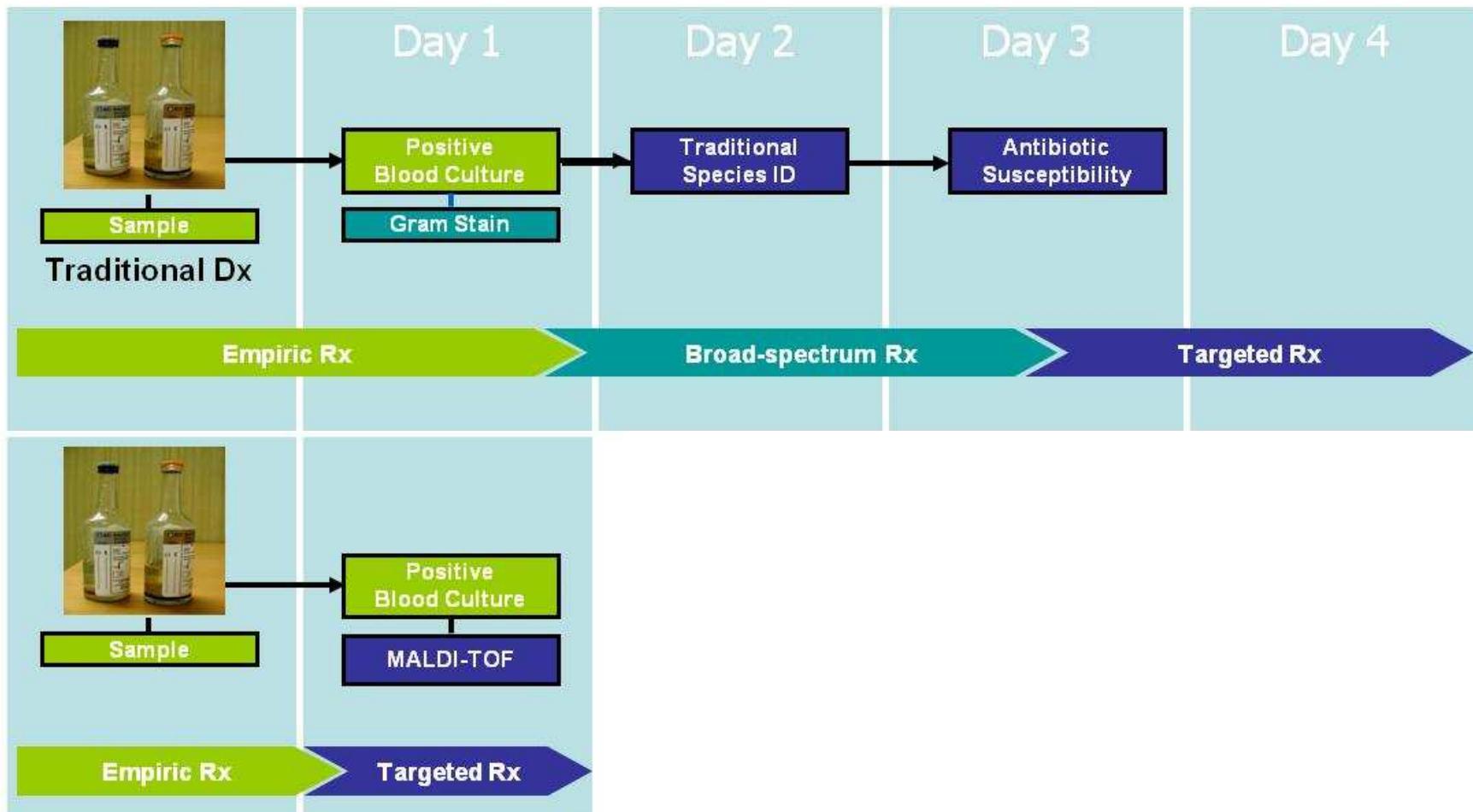
MALDI/MS

New microbiology workflows



Graph adapted from AdvanDx

New microbiology workflows



Graph adapted from AdvanDx

Conclusions

- MS fournit une ID rapide (8h de moins en moyenne), moins cher (environ \$5 US/échantillon) et au moins aussi bons résultats que les approches phénotypique classiques
- L'absence de fausses ID est très rassurant mais une validation minutieuse des nouveaux flux de travail est obligatoire
- Intégration de MS dans le flux de routine et le contrôle de la mise à niveau de la base de données sont les défis actuels
- Des schémas plus élaborés (d'autres matrices, typage de souches par homologie de la signature, etc) devraient être explorées



Var-matin

MS IN THE FUTURE?

- ID of an increasing number of pathogens
(improvement of database)**

- ID of yeasts**

- ID of virulence factors (PVL)**

- ID of resistance (β -lactamase? MRSA?)**

- ID of anaerobes (Actino)**

MALDI-TOF MS FOR YEASTS?

JOURNAL OF CLINICAL MICROBIOLOGY, Sept. 2009, p. 2912–2917

0095-1137/09/\$08.00+0 doi:10.1128/JCM.00389-09

Copyright © 2009, American Society for Microbiology. All Rights Reserved.

Vol. 47, No. 9

Matrix-Assisted Laser Desorption Ionization–Time of Flight Mass Spectrometry for Fast and Reliable Identification of Clinical Yeast Isolates[▽]

G. Marklein,¹ M. Josten,¹ U. Klanke,¹ E. Müller,¹ R. Horré,³ T. Maier,² T. Wenzel,²
M. Kostrzewa,² G. Bierbaum,¹ A. Hoerauf,¹ and H.-G. Sahl^{1*}

*Institute of Medical Microbiology, Immunology and Parasitology, Rheinische Friedrich-Wilhelms-Universität,
University of Bonn, Bonn, Germany¹; Bruker Daltonik GmbH, Bremen, Germany²; and
Federal Institute for Drugs and Medical Devices (BfArM), Bonn, Germany³*

Received 20 February 2009/Returned for modification 9 April 2009/Accepted 22 June 2009

MALDI TOF MS FOR VIRULENCE



Contents lists available at ScienceDirect

International Journal of Antimicrobial Agents

ELSEVIER

journal homepage: <http://www.elsevier.com/locate/ijantimicag>



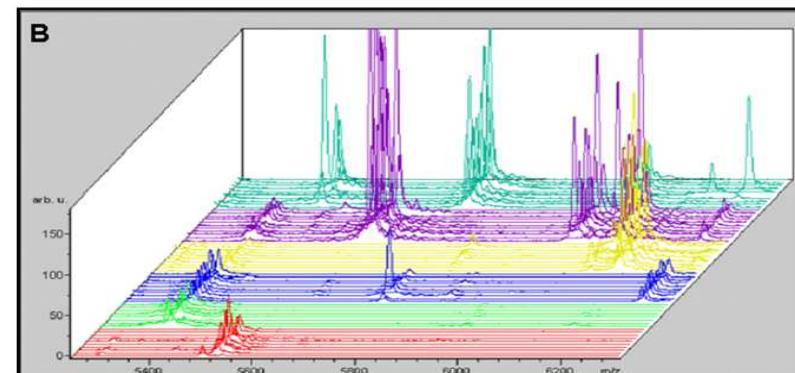
Short communication

MALDI-TOF-MS for rapid detection of staphylococcal Panton–Valentine leukocidin

Fadi Bittar^a, Zoulikha Ouchenane^b, Farida Smati^b, Didier Raoult^a, Jean-Marc Rolain^{a,*}

^a Unité de Recherche sur les Maladies Infectieuses et Tropicales Emergentes (URMITE), CNRS-IRD, UMR 6236, Faculté de Médecine et de Pharmacie, Université de la Méditerranée, 27 Bd Jean Moulin, 13385 Marseille Cedex 05, France

^b Laboratoire de Bactériologie, Hôpital Militaire Universitaire de Constantine, Algeria



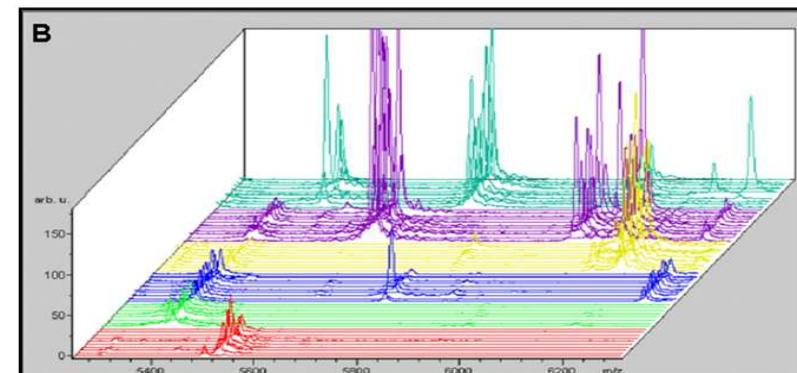
MALDI-TOF MS FOR AB RESISTANCE?

Anal Bioanal Chem (2007) 389:1633–1638
DOI 10.1007/s00216-007-1558-7

TECHNICAL NOTE

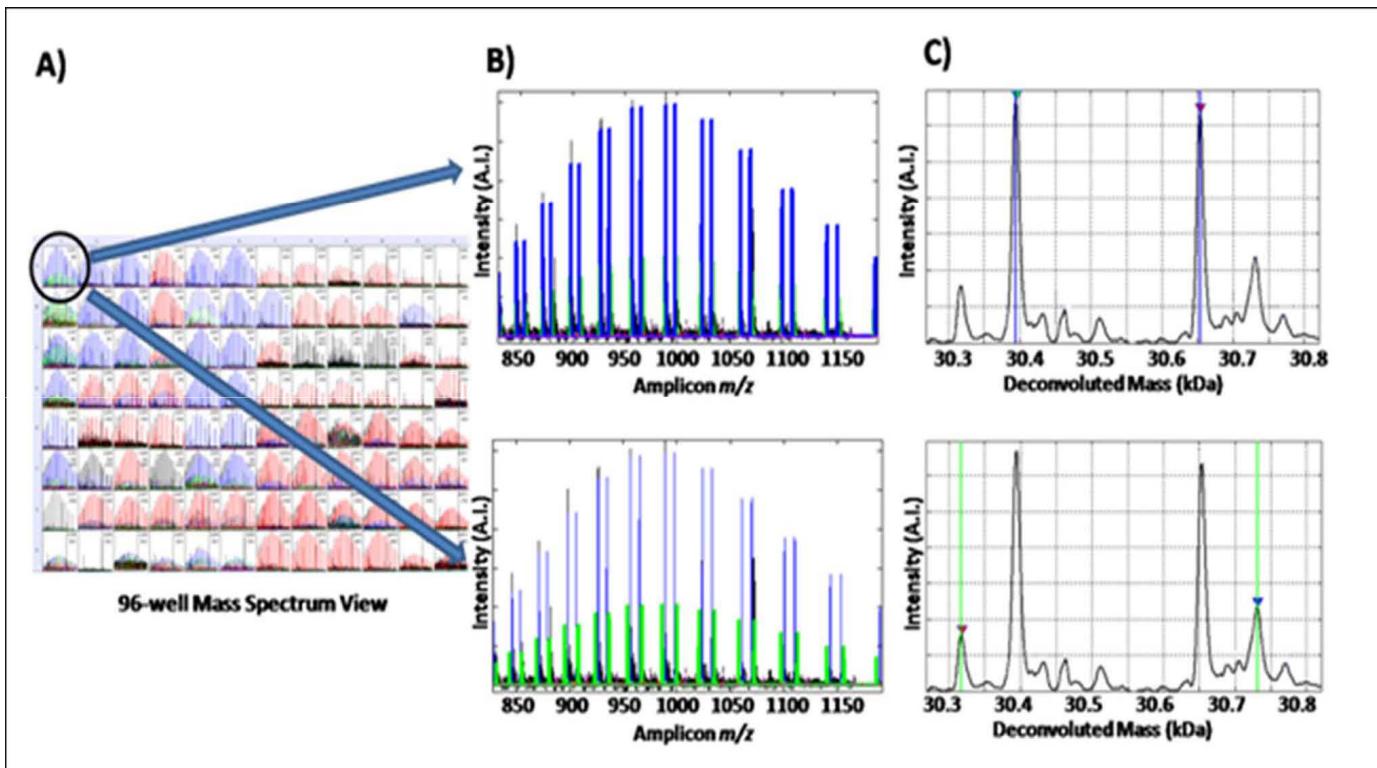
Discrimination between wild-type and ampicillin-resistant *Escherichia coli* by matrix-assisted laser desorption/ionization time-of-flight mass spectrometry

Johanna E. Camara · Faith A. Hays



PCR/ESI-MS

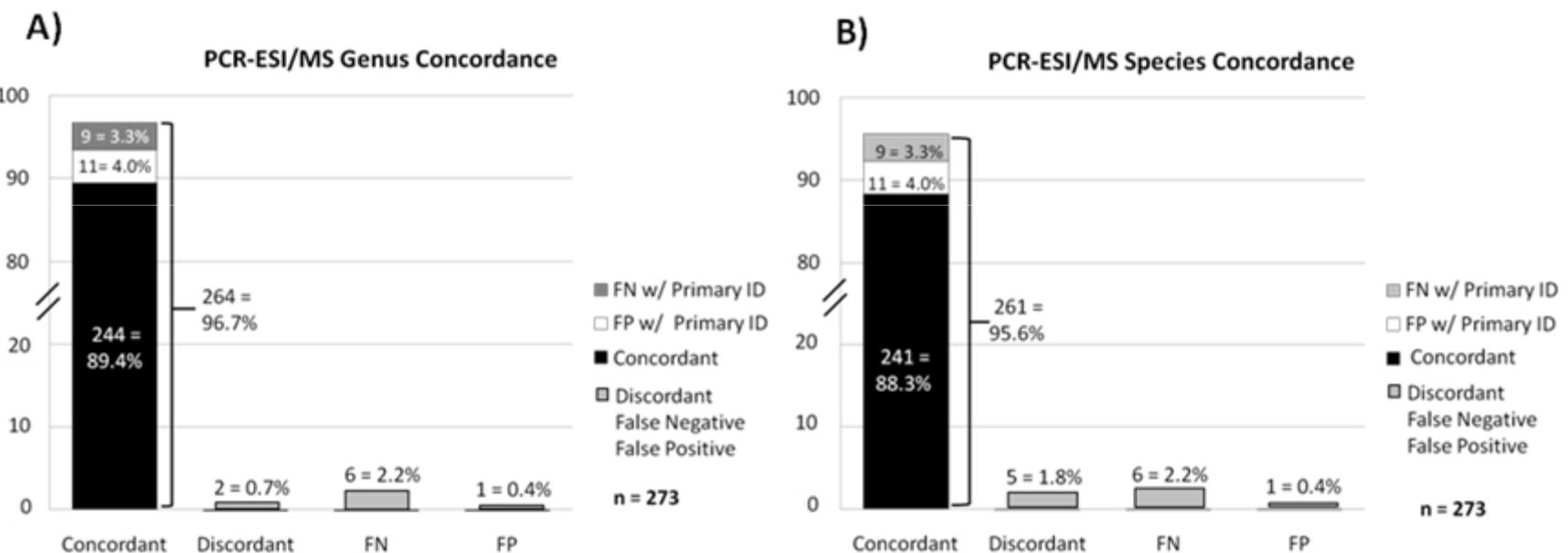
(Electrospray Ionization Mass Spectrometry)



Erin J. Kaleta *et al*, JCM 2010

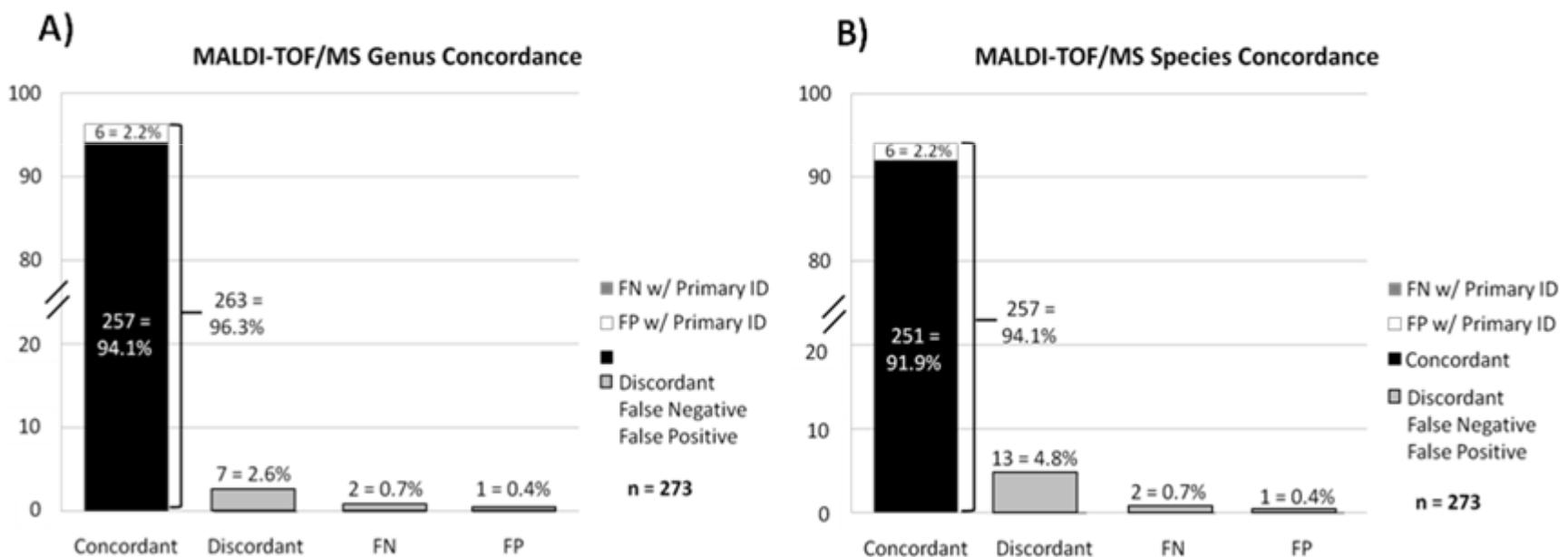
PCR/ESI-MS

(Electrospray Ionization Mass Spectrometry)



PCR/ESI-MS

(Electrospray Ionization Mass Spectrometry)



Merci