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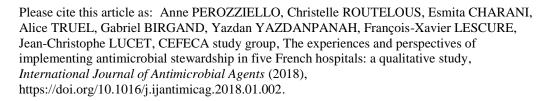
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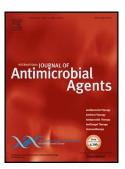
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- 1 The experiences and perspectives of implementing antimicrobial stewardship in five
- 2 French hospitals: a qualitative study

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33	Highlights
34	• The implementation of Antimicrobial Stewardship Program (ASP) depends on
35	organisational, structural and cultural context
36	• We conducted a qualitative study based in five large French hospitals, where ASP were
37	mainly driven by infectious diseases specialists
38	• In this context, with lack of appropriate human and information technology resources,
39	ASP leaders chose adaptive responses and a non-confrontational approach rather than
40	coercive measures
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42	Abstract
43	Objective : To describe current antimicrobial stewardship program (ASP) in France, both at
44	policy level and at local implementation level, and to assess how ASP leaders (ASPL) worked
45	and prioritised their activities.
46	Methods: We conducted a qualitative study based on face-to-face semi-structured
47	interviews with healthcare professionals responsible for ASP across five French hospitals.
48	Five infectious disease specialists and one microbiologist were interviewed between April
49	and June 2016.
50	Results: Stewards had dedicated time to perform ASP activities in two university-affiliated
51	hospitals while in the other hospitals (one university, one general and one semi-private),
52	ASPLs had to balance these activities with clinical practice. Consequently, they had to adapt
53	interventions according to their resources (IT or human). Responding to colleagues'
54	consultation requests formed baseline work. Systematic and pro-active measures allowed
55	for provision of unsolicited counselling, while direct counselling on wards required
56	appropriate staffing. ASPL aimed at increasing clinicians' ability to prescribe adequately and
57	awareness of the unintended consequences of inappropriate use of antibiotics. Thus,
58	persuasive e.g. education measures were preferred to coercive ones. ASPL faced several

- 59 challenges in implementing ASP: overcoming physicians' or units' reluctance, and balancing
- the influence of medical hierarchy and professional boundaries. 60
- Conclusion: Beyond resources constraints, ASPLs' conceptions of their work, as well as 61
- 62 contextual and cultural aspects, led them to adopt a persuasive and collaborative approach
- 63 of counselling. This is the first qualitative study about ASP in France exploring stewards'
- 64 experiences and points of view.
- .microbial re **Keywords**: antimicrobial stewardship, antibiotics, antimicrobial resistance, qualitative study. 65

Introduction

67	The emergence of multidrug resistant bacteria represents a major public health issue, and is
68	addressed by many organisations worldwide[1–4]. According to the Centers for Disease
69	Control and Prevention (CDC), drug-resistant bacteria cause two million illnesses and 23,000
70	deaths annually. Antibiotic resistance is, through the selection pressure mechanism, a
71	consequence of misuse and overuse of antimicrobial agents. To tackle this problem, many
72	international and national recommendations have promoted the implementation of
73	antimicrobial stewardship programs (ASP) in hospitals over the last two decades[5–10]. In
74	France, these programs, when they exist, cover a broad range of organisations[11,12]. In the
75	absence of financial support, each hospital is left to define its own ASP development and
76	implementation.
77	ASP in France relies upon an antibiotics advisor: a practitioner trained in infectious diseases
78	(ID) or in antibiotic treatment, working in close cooperation with pharmacists,
79	microbiologists and infection control practitioners. ASP are most of the time led by ID
80	specialists. Involvement of hospital microbiologists and pharmacists in clinical activities and
81	in "bedside care" is limited. Microbiologists and pharmacists provide advice in their fields of
82	competence and have specific missions. If collaborating, they would form an ASP team.
83	Institutional committees, such as the Anti-infection Agents Committee (CAI), are in charge of
84	local antibiotics policy[10].
85	Our study aimed to describe current ASP in 5 French hospitals, both at policy level and at
86	local implementation level, and to assess how healthcare professionals responsible for ASP
87	are prioritising their activities. Our objective was to understand how the identified antibiotic
88	stewardship program leaders (ASPL) perceived their work and duties in 5 healthcare facilities
89	(HCF), and to discern the factors that influence their strategies and priorities.

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91	This study was part of a larger research project, conducted in randomly selected French
92	HCFs, aiming at describing ASP in France. To validate the study protocol, we carried out a
93	test phase in a convenience sample of 5 HCFs: 2 university-affiliated hospitals, 2 general
94	public hospitals and 1 semi-private hospital. All hospitals are located in Paris or its outskirts.
95	ASPLs were the referring ASP physician or microbiology specialist formally appointed for the
96	hospital. In each setting, ASP leaders were identified and were contacted; all agreed to
97	participate. In April and June 2016, we conducted face-to-face semi-structured interviews
98	with 6 identified stewards: 5 ID physicians and 1 non-physician microbiologist. In one setting
99	ASP leadership was shared by an ID physician and a microbiologist.
100	All interviews were conducted by a researcher (AP), using a semi-structured interview guide,
101	consisting of open-ended questions to explore participants' views and experiences of
102	implementing ASP, including their objectives and priorities, perceived barriers and
103	facilitators to prescribers' uptake, ASP results and prospects. Interview guide is found as
104	supplementary data. All interviews were de-identified and transcribed verbatim. Those
105	interviews were analysed independently by AP and by a sociology lecturer (CR), using an
106	iterative thematic approach[13,14] that resulted in the definition of themes and sub-themes
107	which yielded an analytic framework, used to compare interviews.
108	Ethical approval was obtained from the Hôpitaux Universitaires – Paris Nord Val-de-Seine
109	(HUPNVS) ethical committee (n° 16-018) and all participants signed informed consent prior
110	to the interviews.

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Participants' hospitals and ASP details are presented in table 1. Four hospitals were university-affiliated and one was a semi-private hospital. In all hospitals, the identified ASPL was an ID specialist, coupled with a microbiologist in one. In 2 hospitals, stewards were fulltime dedicated to ASP activities. Microbiology department was involved in ASP activities in all settings, while pharmacists were engaged in 3 hospitals. All hospitals had similar baseline interventions: guidelines on antimicrobial use and management of infectious diseases (local guidelines in 4 hospitals), clinical disease- or speciality-based protocols, and controls or restrictions of sensitive antibiotics, notably carbapenems, third-generation cephalosporins, linezolid or daptomycin. Hospitals all had a dedicated phone number for physicians to call the ASPL. They did not hold consultations after working hours but on-call microbiologists usually took over and provided some advice in the absence of ASPL. Except in one hospital, ASP in its current form started recently (Table 1). However, antibiotic counselling existed previously in the 4 other hospitals in a less formal configuration and with less developed interventions. We conducted interviews with 6 ASPLs, who had a median of 9.5 years of practice (range, 1-15 years). Interviews lasted on average 40 minutes (range, 28-62 mns). The interviews showed several main themes. Physicians' quotations are found in Table 2 [reference TX.X]. Baseline ASP work: response to demands and systematic measures ASP often lack of human resources, as the programs were financially self-supported, with 3 teams without dedicated time to ASP activities. Consequently, ASPLs had to adapt their

activities accordingly, and to prioritise their interventions [T1.1].

Response to solicited consultations from their colleagues from other specialities was
described as the main part of participants' work. Being available and prompt in responding
to their colleagues' demands were considered as a valuable asset, but ASPLs underlined their
difficulties to respond to a growing demand, especially when they took on counselling
besides their clinical work [T1.2].
ASP members also developed systematic interventions, where counselling is initiated by the
stewards, based on laboratory results (e.g. positive blood cultures or multidrug resistant
bacteria), alerts from pharmacists (e.g. controls of antibiotics prescriptions or restricted
antibiotics), or other measures such as systematic control of treatment reassessment at day
3. Using this information, ASPLs contacted prescribers to discuss the clinical situation, and to
offer their help to define treatment or diagnostic strategies. Stewards judged this approach
as "interventionist" as they provide counselling without physicians' request. When ASPLs did
not have dedicated time, they gave priority to remote counselling to contact physicians, over
the phone or with notes in medical files, while bedside consultations were restricted to
complex cases. In the other two HCFs, where ASPLs were dedicated full-time to the ASP,
they chose a more direct and transverse approach: they did rounds in wards, met with
physicians, examined patients, or joined staff meetings. They also had more time to provide
"bedside teaching" to junior doctors. This approach, based on routine counselling, helps to
build cooperative habits between ASPL and ward physicians.
ASPL' approach of counselling: empathic, collaborative and persuasive
Stewards focused more on increasing physicians' capacity to prescribe adequately than on
restricting them in their practices. Participants pointed out that their role was not to control
all antibiotic prescriptions: "It's not my job to control all antibiotic prescriptions". They did

160	not want to be seen as the "antibiotics police", inspecting their colleagues' practices: "I can't
161	verify all antibiotic prescriptions; we can focus on certain units but we must be careful not to
162	discriminate or stigmatise people".
163	ASPLs relied on educating and raising physicians' awareness of the importance of
164	appropriate antimicrobials use and resistance mechanisms to foster new prescribing habits,
165	especially among junior doctors. Every interaction was seen as an opportunity to transfer
166	knowledge around clinical situations [T2.1]. ASPLs were convinced that discussion and direct
167	observation are an essential component of junior doctors' training. Participating in staff
168	meetings with clinicians provide opportunities to discuss directly with physicians, share
169	experience and get the message across about appropriate use of antibiotics [T2.2].
170	Conversely, coercive measures were not regarded as a solution to promote appropriate
171	antimicrobials use. In two hospitals, prescriptions of several antibiotics were "restricted"
172	through software used to prescribe. However, ASPLs were convinced that physicians can 'get
173	round' these restrictions [T2.3].
174	ASPLs tried to be understanding while providing counselling. For instance, when a prescriber
175	disagreed with their advice or recommendations, rather than being authoritarian, ASPL
176	chose to discuss the situation with the ward clinician. Stewards preferred negotiation to
177	confrontation [T3.1]. However, in order to maintain good relations with their colleagues,
178	they admitted conceding non-compliant prescriptions in some cases, so as not to
179	compromise future requests [T3.2].
180	AS were cautious not to put physicians in a difficult or uncomfortable position. Instead, as
181	clinicians themselves, they empathised: they understood physicians' anxiety, their
182	willingness to provide appropriate care to their patients, and the inherent responsibility
183	[T3.3].

Stewards' attitude was also considered decisive for ASP interventions: they did not consider
themselves as the "experts" and the "ones who know" [T4.1]. They respected their peers'
work and clinical judgement: "I'm not going to interfere in their work; at some point, you
have to understand and trust them".
Tailored interventions to adapt to units and physicians' needs and expectations
AS adapted their interventions to units' context and organisation, often described by
departments: medical, surgical, and intensive care units (ICU). They considered the medical
environment of the patient when prioritising their interventions [T5.1]. For instance, in
medical wards, physicians would have considered clinical aspects, requested complementary
exams, and would describe accurately the situation to the AS when asking for advice. While
in surgery wards, AS preferred to go and examine the patient, or to discuss with the non-
medical staff, as surgeons spend most of their time in the operating room and are not too
comfortable with medical aspects of patient care. AS considered that usually, there was no
need to go to or focus on ICU units, as physicians there were as knowledgeable and capable
as them to deal with infectious situations. These wards were not a priority of ASP
interventions: "we are not going to progress by being picky with ID or ICU physicians, they
are well-informed".
One other asset of ASPLs strategies was to adapt their responses to clinicians' demands.
Their inputs varied according to what physicians expected from them. We identified three
levels of involvement:
Help physicians in the decision making process: ASPLs have specific knowledge in ID and
microbiology. One of their roles was to help or to comfort clinicians in their decision,
while promoting appropriate antibiotic use. This was the most frequent reason for

208	physicians' calls. In those situations, prescribers seemed to follow easily stewards'
209	recommendations as they conceded that ID specialists had more advanced knowledge
210	regarding bacteriology and antibiotics properties [T5.2].
211	Reduce physician's uncertainty: When they faced a complex case, ward physicians
212	sought ID specialists' expertise to overcome their own limits or doubts. ASPLs'
213	recommendations and guidance reassured the physician in charge. Stewards were not
214	directly in charge of the patient so they could help the physician by giving an "external"
215	opinion. One way for ID specialists to respond to physicians' uncertainty was to engage
216	their own responsibility in the decision, e.g. written recommendations in patient's chart
217	[T5.3].
218	Delegation of responsibility: In some cases, prescribers delegated antibiotics prescribing
219	and infectious situations management to the stewards. This was more frequent in
220	surgery wards, where surgeons accepted to share medical care of their patients with
221	other physicians. Delegation was a pragmatic solution that benefit to both parties,
222	improving patients' care and optimising antibiotics prescribing. However, this was not a
223	path all participants wanted to take, as it requires time and results in blurred boundaries
224	between physicians and ASPLs' tasks and responsibilities [T5.4].
225	
226	ASP challenges
227	Several challenges were reported. ASPLs' interventions were driven by physicians' demands:
228	they sought ID specialists' expertise and their updated knowledge to adapt antibiotic
229	therapy to bacteriological results or clinical situation, but less often to discuss diagnostic
230	strategies [T6.1]. They were not always inclined to involve stewards in patient care. ASPLs

wished they had been soli	cited more often for their clinical skills and knowledge, and to be
sometimes confused with	microbiologists or considered as "antibiogram interpreters".
The physician in charge of	the patient care remains the person legally responsible for the
patient. The ASPLs had to	respect clinical jurisdiction over patient: they made advisory
recommendations, but did	d not interfere with physicians' ultimate decisions regarding their
patients [T6.2]: "The phys.	ician in charge makes the choice, I only provide advice".
As a result, in all hospitals	, ASPLs did not have access to certain units that did not solicit
them and were not recept	tive to unsolicited advice. Moreover, the participants in this study
did not report any strateg	ies to overcome reluctance from individual physicians or from
entire units.	
Hierarchical influence was	another main challenge inherent to ASP interventions. Prescribing
habits are influenced by the	ne practices of peers or superiors. ASPLs' recommendations
competed sometimes with	n wards' practices or senior doctors' instructions [T6.3]. In case of
discrepancies, AS acknowl	edged junior doctors' difficulty to follow their advice, as it would
mean to argue the decisio	n of a superior. In their daily activity, stewards communicated
mostly with junior doctors	and residents, as they were the ones present in the ward and who
write the prescriptions. Ye	et, the lack of direct communication between senior doctors and
ASPLs was considered as a	an obstacle, as they could not explain their opinion and justify why
they had made such recor	nmendation [T6.4].

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Discussion

252	Previous studies described ASP activities in French hospitals[11,12], but to our knowledge,
253	this is the first qualitative study about ASP ever performed in France.
254	This work suggests that ASPLs had to adapt their interventions to the available resources,
255	including human and information technology (IT) resources and time dedicated to
256	counselling activity. Stewardship activities consisted mainly in ID physicians-stewards
257	counselling colleagues about antibiotic use or infectious diseases management. Systematic
258	advice based on microbiological results (bacteraemia) or pharmacists' controls depended on
259	available IT resources and local collaborations. Despite ASP are mandatory and endorsed by
260	hospital's authorities[15], the support of the hospital governance was somehow
261	"theoretical" and usually did not ensure appropriate human, financial and IT resources for
262	ASP teams. Consequently, in the absence of detailed regulatory framework and adequate
263	resources, ASP organisation and activities were often non-formalised.
264	Beyond these constraints, ASPLs' conceptions of counselling led them to choose a persuasive
265	and collaborative approach over a more restrictive one. ASP was regarded as "an advisory
266	service" [16] opposed to a policing body. Participants did not describe their role as
267	controlling or reviewing all antibiotic prescriptions. Conversely to what is described in the
268	literature and suggested by international authorities[7,17–19], ASP in our sample included
269	few restrictive measures, such as prior authorisation or expert approval, therapeutic
270	substitution or automatic stop orders. Instead, ASP teams gave priority to educative
271	strategies: formal educational sessions (essentially towards junior doctors) and informal
272	knowledge transfer through interactions with prescribers. When ASPLs had dedicated time,
273	they were able to provide bedside teaching. Observations and discussions with ID-specialists
274	represent an informal form of training for junior or less experienced doctors[20,21].

Educational strategies take time but are associated with higher acceptance and long-term
effect.
Several reasons may explain why ASP in France are based on persuasive and collaborative
approaches of counselling: 1) lack of resources (IT and human resources) prevents
considering restrictive measures, such pre-authorisation or automatic stop orders. These
restrictive interventions required computerised surveillance and stewards' availabilities to
approve antimicrobial prescriptions; 2) cultural aspects: according to Hofstede's model of
cultural dimensions, France had a high degree of power distance, indicating a high level of
hierarchy and an unequally distributed power, as well as a high level of individualism[22]. In
this context, collaboration between doctors seems less straightforward and physicians'
clinical autonomy probably stronger. Restricting clinicians in their intentions and actions
would go against this fundamental principle of medical practice and could be considered a
hindrance to their work. Professional hierarchic differences would explain why, despite the
lack of human resources and ID physicians-stewards' difficulties to achieve all ASP
objectives, microbiologists and pharmacists' involvement in antibiotics counselling and other
ASP primary missions may be insufficient.
ASPLs also emphasised the necessity of building collegial relationships with wards clinicians.
Several studies concluded that strategies based on delivering technical advice or
disseminating guidelines were not sufficient to improve antibiotics prescribing behaviours.
Conversely, investing in interprofessional relationships and effective collaboration were
identified as a key process for ASP and had more sustainable impact[16,23,24]. ASPLs also
adopted a non-confrontational attitude when providing advice, even when physicians
disagreed with them. This non-judgemental attitude may create a "safe environment" for
physicians and the quality of interactions is essential to ensure further requests from

299	physicians[7]. Conversely, Goldstein mentioned the potential disastrous effect of
300	authoritarian approaches or overruling of physicians' clinical judgement[25].
301	ASPLs believed that adapting their strategies to the local context increased the efficacy of
302	their interventions and physicians' uptake. Cortoos underlined the need to differentiate
303	between specialties when deploying ASP interventions as prescribers were a heterogeneous
304	population, with different attitudes and expectations[26]. For instance, surgeons shared
305	more easily decisions regarding patient's medical care. The absence of competition between
306	surgeons and ASPLs qualification and tasks make makes this type of cooperation easier.
307	Stewards considered ICU physicians more competent than in other units to manage
308	antimicrobial, and felt that their input would be more beneficial in other wards. In addition,
309	ICU physicians may argue patients' severity to justify the use of broad-spectrum antibiotics.
310	The situation is more complex in medical units where physicians may refrain from involving
311	ID specialists in patients care. Their reservations could be explained by: the overlap of
312	medical knowledge and common training among medical specialties, the sense of ownership
313	of clinical decision-making, the high specialisation of hospital physicians and patients'
314	specificities (comorbidities, severe conditions)[23,27,28]. Physicians frequently consider they
315	can manage infectious diseases by themselves, especially the ones within their own
316	speciality, and sometimes fail to acknowledge the input of a specialist, especially regarding
317	diagnostic issues.
318	This persuasive and "non-constraining" approach of ASP had its drawbacks and ASPLs had to
319	face several challenges: to balance the influence of medical hierarchy and professional
320	boundaries, and to overcome physicians and units' reluctance.
321	In hospitals, ASPLs had a role as consultants on antimicrobial treatments or infections, while
322	ward physicians are medically and legally in charge of patients. Moreover, Oh explained that

in teaching hospitals, the hierarchical structuring of care delivery forms another obstacle for
consultants, as interns and residents receive orders from their superiors which they feel
compelled to carry out[28]. Many studies described how hierarchical influence and
prescribing etiquette weigh on junior and even more experienced doctors, leading them to
adopt usual practices that could not be questioned by their colleagues[29–32]. Several
authors[23,29,30] described the role of senior doctors in knowledge transfer in medicine and
how antibiotic prescribing habits, among other things, are passing from seniors to juniors.
This study questioned ASP strategies targeting interns, and suggested the need to increase
direct communication with attending physicians. ASP interventions could also consider
addressing potentially outdated information or gaps in practising physicians' knowledge[33].
When confronted to reluctant units, participants did not develop strategies to overcome
physicians' resistance. As sometimes newly implemented, ASPLs' priority was first to get
physicians' uptake and confidence. Yet, they need to develop solutions to overcome barriers.
Several strategies could be considered, such as demonstrating positive outcomes in patients
with appropriate antibiotics treatment [24,25,34]. A global policy about antibiotics at the
hospital level, supported by administrative and medical heads would also probably
strengthen ASPL's position and help to overcome units' reluctance. The intervention of a
supra-level body may be necessary, such as antimicrobial committees, and the involvement
of hospital authorities may be required[23,35].
Choosing to provide an incentive rather than a more restrictive approach had limitations,
especially to overcome physicians' lack of interest or unwillingness to conform to
recommendations regarding antibiotic prescriptions. This may partly explain the high level of
antibiotic consumptions in French HCFs[36].

This study had several limitations: it was based on a small sample of hospitals (5) and
interviews (6), which were not geographically representative of all settings. Furthermore,
this study did not reflect the work and perceptions of all potential members of ASP, such as
microbiologists and pharmacists. This would explain why stewards reported few controls and
restrictive measures, as these tasks usually fall to pharmacists. Our next work will explore all
stewards' roles and perceptions, in a larger sample of hospitals.
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353	Conclusion
354	Our findings highlight how ASP had to adapt to organisational, structural and cultural
355	context. ASP consisted mainly in an ID counselling activity rather than in a stewardship
356	program, managed by a multidisciplinary team. ASPLs chose adaptive responses and non-
357	confrontational approach that would make interventions easier to implement and less
358	subject to opposition from prescribers. ASP policy must be formally endorsed and supported
359	by hospital administration to overcome acceptance barriers. The lack of appropriate human
360	and IT resources also affects ASPLs' interventions and limits systematic measures. Current
361	ASP may not meet their goals aiming at decreasing antimicrobial resistance and use.
362	
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375	
376	Supplementary data: Interview guide on stewards' work and perceptions

377	Refe	erences
378	[1]	World Health Organization - WHO. Global action plan: on antimicrobial resistance 2015.
379	[2]	Carlet J, Le Coz P. Together, let's save antibiotics. Proposals of the special working
380		group for keeping antibiotics effective 2015.
381	[3]	Action plan against the rising threats from Antimicrobial Resistance. Communication
382		from the Commission to the European Parliament and the Council. 2011.
383	[4]	Centers for Disease Control and Prevention - CDC. Antibiotic resistance threats in the
384		United States, 2013.
385	[5]	Barlam TF, Cosgrove SE, Abbo LM, MacDougall C, Schuetz AN, Septimus EJ, et al.
386		Implementing an Antibiotic Stewardship Program: Guidelines by the Infectious Diseases
387		Society of America and the Society for Healthcare Epidemiology of America. Clin Infect
388		Dis 2016;62:e51-77.
389	[6]	Pollack LA, Srinivasan A. Core elements of hospital antibiotic stewardship programs
390		from the Centers for Disease Control and Prevention. Clin Infect Dis 2014;59 Suppl
391		3:S97-100.
392	[7]	Dellit TH, Owens RC, McGowan JE, Gerding DN, Weinstein RA, Burke JP, et al. Infectious
393		Diseases Society of America and the Society for Healthcare Epidemiology of America
394		guidelines for developing an institutional program to enhance antimicrobial
395		stewardship. Clin Infect Dis 2007;44:159–77.
396	[8]	European Centre for Disease Prevention and Control. Proposals for EU guidelines on
397		the prudent use of antimicrobials in humans. Stockholm: ECDC; 2017
398	[9]	MINISTÈRE DES AFFAIRES SOCIALES, DE LA SANTÉ, ET DES DROITS DES FEMMES,
399		Direction générale de l'offre de soins. Instruction DGS/RI1/DGOS/PF2/DGCS n° 2015-

400		212 du 19 juin 2015 relative à la mise en œuvre de la lutte contre l'antibiorésistance
401		sous la responsabilité des agences régionales de santé 2015.
402	[10]	Haute autorité de Santé - HAS. Antibiotic therapy and prevention of bacterial resistance
403		- Guidelines - April 2008.
404	[11]	Le Coz P, Carlet J, Roblot F, Pulcini C. Human resources needed to perform
405		antimicrobial stewardship teams' activities in French hospitals. Med Mal Infect
406		2016;46:200–6.
407	[12]	Perut V, Aumaître H, Pichard E, Patey O, Andre P, Welker Y, et al. Transversal infectious
408		disease activity in French hospitals. Médecine Mal Infect 2017;47:50–7.
409	[13]	Fereday J, Muir-Cochrane E. Demonstrating rigor using thematic analysis: A hybrid
410		approach of inductive and deductive coding and theme development. Int J Qual
411		Methods 2006;5:80–92.
412	[14]	Guest G, Namey EE, Mitchell ML. Collecting Qualitative Data: A Field Manual for
413		Applied Research. SAGE Publications; 2012.
414	[15]	Décret n° 2013-841 du 20 septembre 2013 modifiant les dispositions relatives à la
415		commission médicale d'établissement et aux transformations des établissements
416		publics de santé et à la politique du médicament dans les établissements de santé
417	[16]	Pakyz AL, Moczygemba LR, VanderWielen LM, Edmond MB, Stevens MP, Kuzel AJ.
418		Facilitators and barriers to implementing antimicrobial stewardship strategies: Results
419		from a qualitative study. Am J Infect Control 2014;42:S257-263.
420	[17]	Pollack LA, Srinivasan A. Core elements of hospital antibiotic stewardship programs
421		from the Centers for Disease Control and Prevention. Clin Infect Dis 2014;59 Suppl
422		3:S97-100.

123	[18]	National institute for Health and Care Excellence - NICE. Antimicrobial stewardship.
124		Quality standards, 2016.
125	[19]	National Quality Partners Playbook: Antibiotic Stewardship in Acute Care, 2016.
126	[20]	Skodvin B, Aase K, Charani E, Holmes A, Smith I. An antimicrobial stewardship program
127		initiative: a qualitative study on prescribing practices among hospital doctors.
128		Antimicrob Resist Infect Control 2015;4:24.
129	[21]	Noble C, Brazil V, Teasdale T, Forbes M, Billett S. Developing junior doctors' prescribing
130		practices through collaborative practice: Sustaining and transforming the practice of
131		communities. J Interprof Care 2017;31:263–72.
132	[22]	Deschepper R, Grigoryan L, Lundborg CS, Hofstede G, Cohen J, Kelen GVD, et al. Are
133		cultural dimensions relevant for explaining cross-national differences in antibiotic use
134		in Europe? BMC Health Serv Res 2008;8:123.
135	[23]	Broom J, Broom A, Plage S, Adams K, Post JJ. Barriers to uptake of antimicrobial advice
136		in a UK hospital: a qualitative study. J Hosp Infect 2016;93:418–22.
137	[24]	Jeffs L, Thampi N, Maione M, Steinberg M, Morris AM, Bell CM. A Qualitative Analysis
138		of Implementation of Antimicrobial Stewardship at 3 Academic Hospitals:
139		Understanding the Key Influences on Success. Can J Hosp Pharm 2015;68:395–400.
140	[25]	Goldstein EJC, Goff DA, Reeve W, Naumovski S, Epson E, Zenilman J, et al. Approaches
141		to Modifying the Behavior of Clinicians Who Are Noncompliant With Antimicrobial
142		Stewardship Program Guidelines. Clin Infect Dis 2016;63:532–8.
143	[26]	Cortoos P-J, De Witte K, Peetermans WE, Simoens S, Laekeman G. Opposing
144		expectations and suboptimal use of a local antibiotic hospital guideline: a qualitative
145		study. J Antimicrob Chemother 2008;62:189–95.

446	[27]	Pavese P, Sellier E, Laborde L, Gennai S, Stahl J-P, François P. Requesting physicians'
447		experiences regarding infectious disease consultations. BMC Infect Dis 2011;11:62.
448	[28]	Oh H. Hospital consultations and jurisdiction over patients: consequences for the
449		medical profession. Sociol Health Illn 2014;36:580–95.
450	[29]	Charani E, Castro-Sanchez E, Sevdalis N, Kyratsis Y, Drumright L, Shah N, et al.
451		Understanding the determinants of antimicrobial prescribing within hospitals: the role
452		of "prescribing etiquette." Clin Infect Dis 2013;57:188–96.
453	[30]	Lewis PJ, Tully MP. Uncomfortable prescribing decisions in hospitals: the impact of
454		teamwork. J R Soc Med 2009;102:481–8.
455	[31]	Livorsi D, Comer A, Matthias MS, Perencevich EN, Bair MJ. Factors Influencing
456		Antibiotic-Prescribing Decisions Among Inpatient Physicians: A Qualitative
457		Investigation. Infect Control Hosp Epidemiol 2015;36:1065–72.
458	[32]	Broom A, Broom J, Kirby E. Cultures of resistance? A Bourdieusian analysis of doctors'
459		antibiotic prescribing. Soc Sci Med 1982 2014;110:81–8.
460	[33]	Rocha-Pereira N, Castro Sanchez E, Nathwani D. How can multi-professional education
461		support better stewardship? Infect Dis Rep 2017;9.
462	[34]	Goff DA, Kullar R, Bauer KA, File TM. Eight Habits of Highly Effective Antimicrobial
463		Stewardship Programs to Meet the Joint Commission Standards for Hospitals. Clin
464		Infect Dis 2017;64:1134–9.
465	[35]	Dyar OJ, Tebano G, Pulcini C, ESGAP (ESCMID Study Group for Antimicrobial
466		stewardshiP). Managing responsible antimicrobial use: perspectives across the
467		healthcare system. Clin Microbiol Infect 2017.
468	[36]	European Centre for Disease Prevention and Control. Geographical distribution of
469		antimicrobial consumption 2017.

Table 1: Hospitals characteristics and ASP details

Hospital	Н1	Н2	Н3	Н4	Н5	
Number of acute care beds	424	323	423	848	385	
	ASP organisation					
Existence of ASP (current form)	10 years	6 months	1 year	1.5 years	6 months	
Stewards	1 ID specialist 1 bacteriologist	1 ID specialist	3 ID specialists (rotations)	1 ID specialist (senior) 1 resident	2 ID specialists (senior, rotations) 1 resident	
Dedicated ASP time (full- time)	No	No	No	Yes	Yes	
Microbiology involved*	Yes	Yes	Yes	Yes	Yes	
Pharmacy involved*	Yes	No	Yes	No	Yes	
ASP measures and interventions						
Dedicated ASP phone number or direct line	Yes	Yes	Yes	Yes	Yes	
Mode of consultation	Phone (90%)	50% phone 50% bedside advice	Phone Beside consultations for complex cases	Bedside consultations (80%) Phone for simple questions (20%)	Bedside consultations (70%) Phone for simple questions	
Systematic counselling	Laboratory results (daily monitoring) Pharmacists alerts	Controls of treatment reassessment at day 3 Alerts from microbiology and pharmacy	Case reviews based on laboratory results or alerts from pharmacy	Laboratory results (daily monitoring)	Laboratory results (daily monitoring) Alerts from pharmacy	
ID on wards	No – or complex cases (ID specialist)	Yes – solicited consultations	Yes - solicited consultations	Yes – routine wards rounds, solicited counselling	Yes – solicited and unsolicited counselling	

and staff

and staff

* Formally involved in ASP team and active in ASP missions



Table 2: Quotes from participants, by key themes

Key themes	Reference in paragraph	Quotes
Baseline AS work [T1]	Prioritising interventions and systematic	"I'm with the microbiology resident, who is dedicated to the counselling activity, and
	measures	so we call in systematically all positive blood cultures, but also surgical sample
		collections, so it's mainly done over the phone. I sometimes go to the units for
	Because AS time was sometimes	complex cases, but we try as I don't work full time for the counselling activity, I
	constrained, they favoured a "quantitative"	only have 2 or 3 hours a day, so we tend to provide advice over the phone rather
	approach of counselling (T1.1)	than in person"
	Combining counselling and clinical activity	"Some days, it's complicated. For instance, I hold consultations on Friday
	(T1.2)	afternoons, and it is a busy time for antibiotic counselling. I have calls from my
		colleagues during my consults"
		"I give advice but it is difficult to follow up on the case, to go back 48h later to see if
		they still need help. I don't have the time to do that"
Teach rather than	Every interaction is the opportunity to	"Every time I go to the wards to meet doctors, I explain why we choose this
control [T2]	transfer or update knowledge	antibiotic; we don't just say: "You have to prescribe Vancomycin". We constantly try
	Bedside teaching (T2.1)	to teach them, it's really bedside education"
	× ("That's why I want to organise a staff meeting, because I'm sure that when you
	Staff meeting (T2.2)	have several cases, we discuss them together, and over time we discuss similar
		clinical cases again and again, and after a while they understand and they change
		their prescribing habits"
	Coercive methods: physicians can get	"I think it's more important to educate people. Because they will always find a way
	around restrictions (T2.3)	to prescribe what they want, to bypass the system. They need to understand why
	X .	they shouldn't prescribe this antibiotic".
A comprehensive	In case of disagreement, AS prefer	"I often discuss with the prescriber "So, we have these options, it's up to you, you
approach [T3]	negotiation to confrontation with the	can start with that, and then do that, you choose". And sometimes, I remind them
	physician in charge (T3.1)	that I'm not the one in charge of the patient, it's not my responsibility, so if they are
		concerned, they can start antibiotics, but I ask them to call me back after 48-72h to
		reassess the situation"

	When the situation is sensitive, they avoid conflicts in order to maintain good relations with the unit (T3.2)	"We don't argue, the idea is not to close off from the service. We know our limits, sometimes we try to talk it over but if we feel that they won't agree and the situation can worsen, instead of risking compromising our future collaboration, we give up, yes".
	As clinician themselves, stewards understand the position their colleagues are in (T3.3)	"Very often, a physician faces a patient who is not doing well, who has a fever, and he thinks of an infection, but he doesn't know exactly where the problem is, it takes him time to understand, he has no bacteriological results, the patient is fragile, he is old, and the physician really wants to treat him with antibiotics. And I understand, I was in this situation once, I do understand!"
Stewards' attitudes [T4]	Non authoritative (T4.1)	"It is important to respect our colleagues. I have met stewards who were commanding, but I think it's just counterproductive" "I think that what helps, is that they see I'm available, I answer the phone, I come to the ward and look at the situation with them, and that I'm not categorical, I give them advice"
Tailored responses to units and physicians' demands and expectations [T5]	Need to adapt interventions to units' organisation and context (T5.1)	"We need to adapt to units. For instance, the patient's follow-up after I gave recommendations: when the patient is in a surgery unit, I come back every day to check on him, while if the patient is in a medical unit, where I know that the physicians are autonomous, I don't need to go back every day, because I know physicians won't like it and will find it intrusive, and there is no point to do so, so I only go and check on the patient every 3 or 4 days, or once a week to see if everything is ok. But really, we need to adapt to the patient's medical context for the follow-up" "In medical wards, I don't need to go and see the patient; I know my colleagues would have examined the patient and they can explain the situation to me. My colleagues in surgical wards I go see them"
	To help or comfort physicians in the decision-making process (T5.2)	"They (the wards physicians) expect us to tell them what are the appropriate antibiotics and dosage, and sometimes they don't even know what kind of antibiotics to use. When they have the laboratory results, they also ask sometimes

		what to do with them. Our job is also to remind them that a positive specimen
		should not automatically lead to a treatment".
		"In many cases, they would have prescribed by themselves, but since I'm here, they
		call and ask for confirmation, sometimes because they have to deal with multidrug-
		resistant bacteria and they need an alternative treatment to carbapenem, or they
		may have questions about dosage. Usually, they think of an infection, they want to
		treat the patient, so they call either to validate with me the indication of the
		treatment or to discuss molecules, doses, the duration of the treatment"
	To address physicians' uncertainty: AS	"It is always easier to have an outside position, to give advice rather than to be the
	engage their personal responsibility (T5.3)	one in charge. When you're in charge of a patient, you're always concerned and
		you want him to get better. So, I think that it is a good thing that another physician
		gives an opinion, because we are less stressed, we don't feel this pressure and
		there is less culpability because "I performed the intervention, so the infection is my
		fault". Of course, I'm also worried when I say: "No, there is no need to treat the
		wound for now" but I have to put my responsibility on the line".
	Delegation of antibiotics prescription and	"Some surgical units delegate the choice of antibiotic, its duration, but we discuss
	infectious diseases management (T5.4)	with them when there are diagnostic issues"
	XX	"In some surgical units, we go and see the nurses that tell us which patients are
		currently being prescribed antibiotics. We do our rounds. Surgeons rely completely
		on us for antibiotics management"
ASP' limitations	Physicians' demands (T6.1)	"Physicians think they don't need our help to make a diagnosis but rather to adapt
	20	antibiotic treatments, its timing, and its regimen. But I think that we can also have
		an input, by examining patients, and proposing a differential diagnosis. In some
		units, physicians tell us "No there is no need for you to examine the patient, if there
		are positive cultures, I'll call you".
		"I consider we have an input when making diagnoses, but they (the ward physicians)
		don't realise it, and they don't think about calling us in this regard. They need us
		when they have specimen results, and they have to choose what drugs to prescribe"
		"This unit, they only call when they have a bacteriological issue, because they are

	not comfortable, they are not sure, but they make few requests to discuss diagnostic
	strategies"
Clinical jurisdiction	"I don't interfere in their work. At some point, you have to trust your colleague the same way they trust you"
The mule of an inference prevails (TC 2)	, , ,
The rule of on-inference prevails (T6.2)	"We try not to be intrusive, I do counselling, I can even say: "If I was you, I would do
	that", but I suggest, I don't give orders! It's important to keep some distance, and
	leave the final decision to the prescriber"
Hierarchical influence	"We discuss a lot with residents, because they are the ones present on the ward,
	and the fellow or the attending, who was not there when we make our
	recommendations, does not agree with us, and so, the intern is going to follow his
Junior doctors follow ultimately their	superior's opinion. I think this situation is quite frequent."
supervisor' instructions (T6.3)	"In 99% of the cases, we discuss with the residents. Not with the attending
	physicians (). Afterwards, if the senior says: "Don't listen to them", you can be sure
	that the intern is not going to listen to us! But in reverse, if he says "Ask the AMS
	team, because they know best, you have to call them", then the interns will do it
Lack of direct communication between AS	willingly"
and attending physicians (T6.4)	"We give advice to the resident, who is going to agree (). But his superior may
0,7	have a different opinion, and then he changes the prescription. It happens when we
	did not talk to the attending directly".

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