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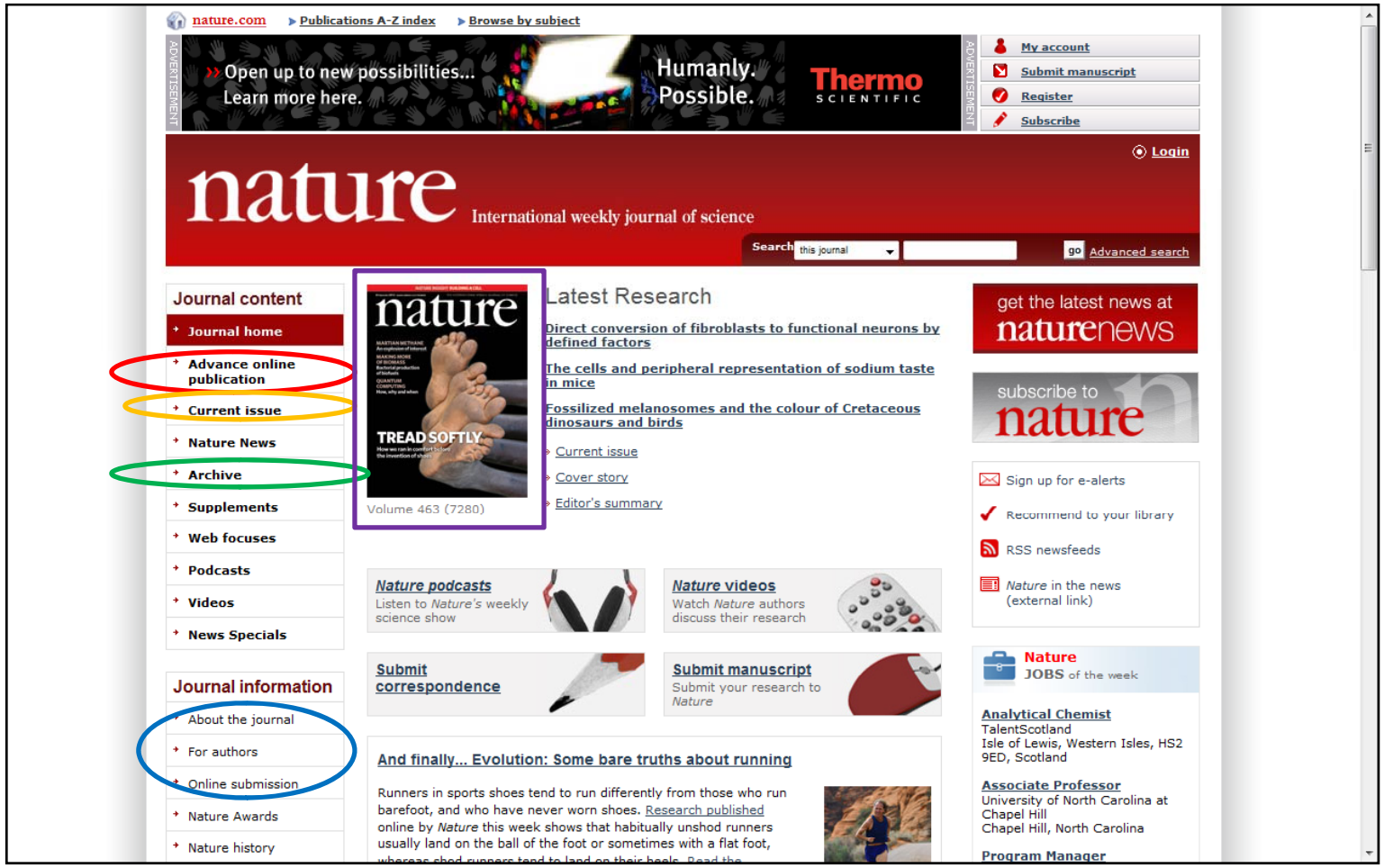

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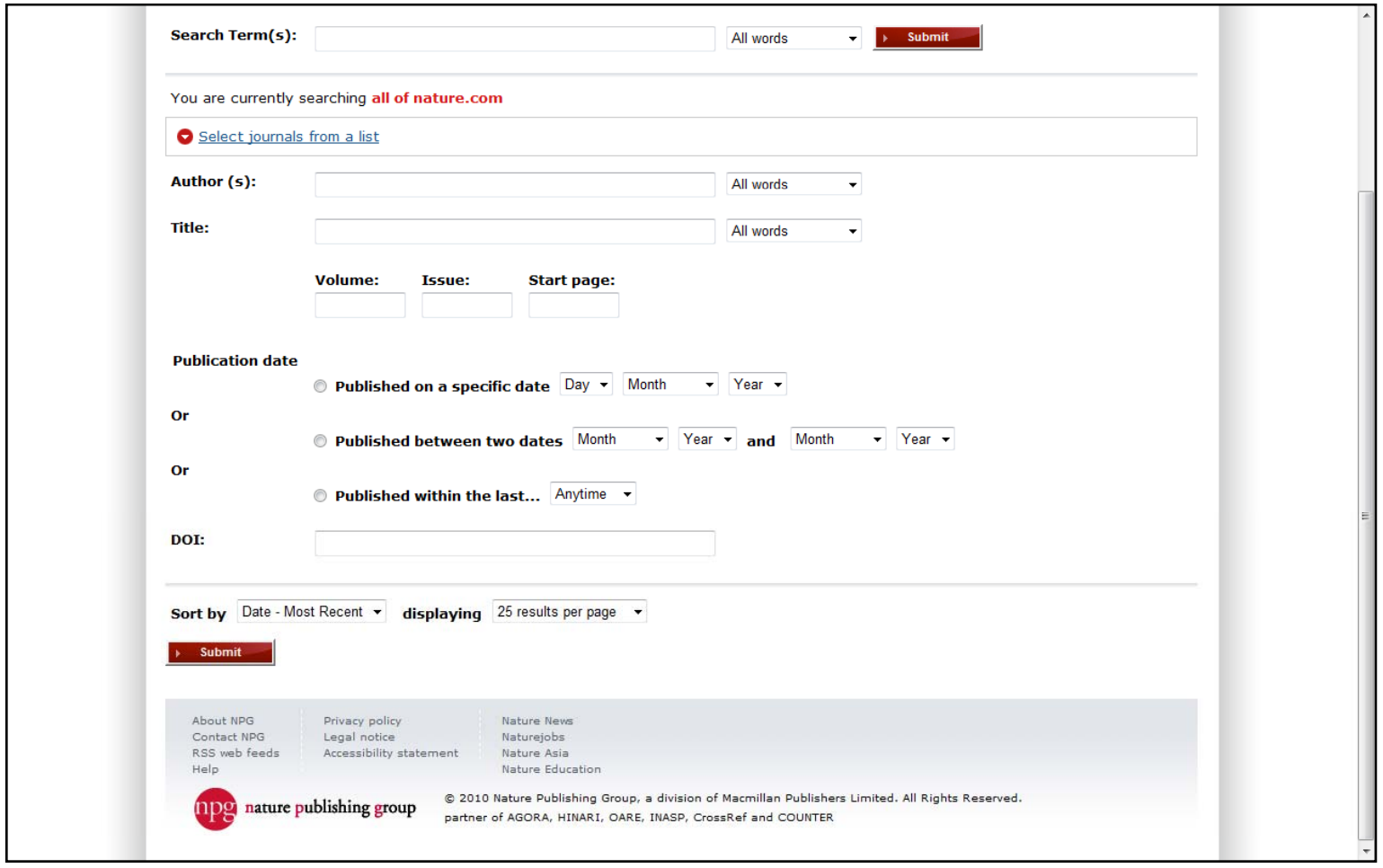

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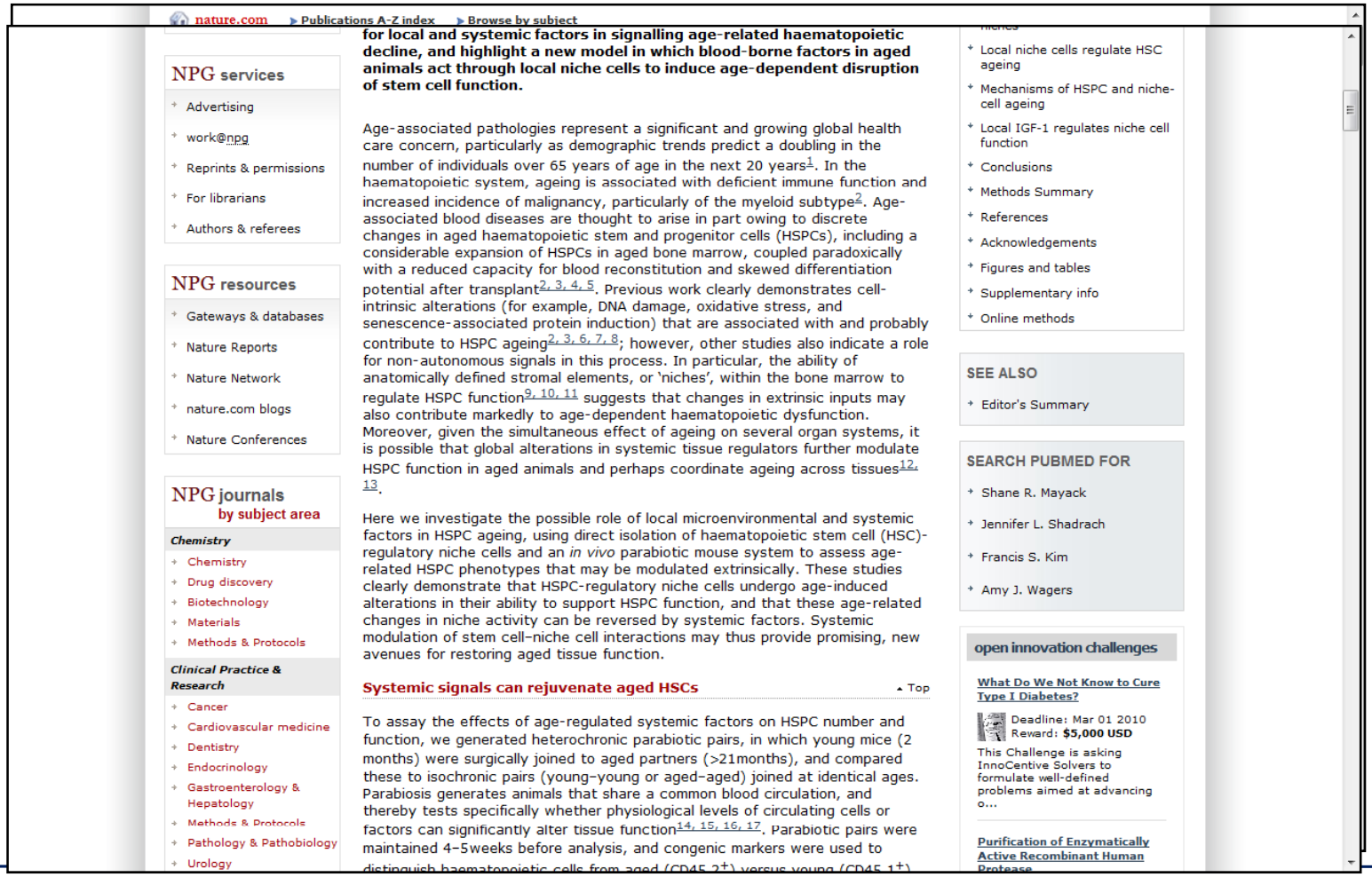

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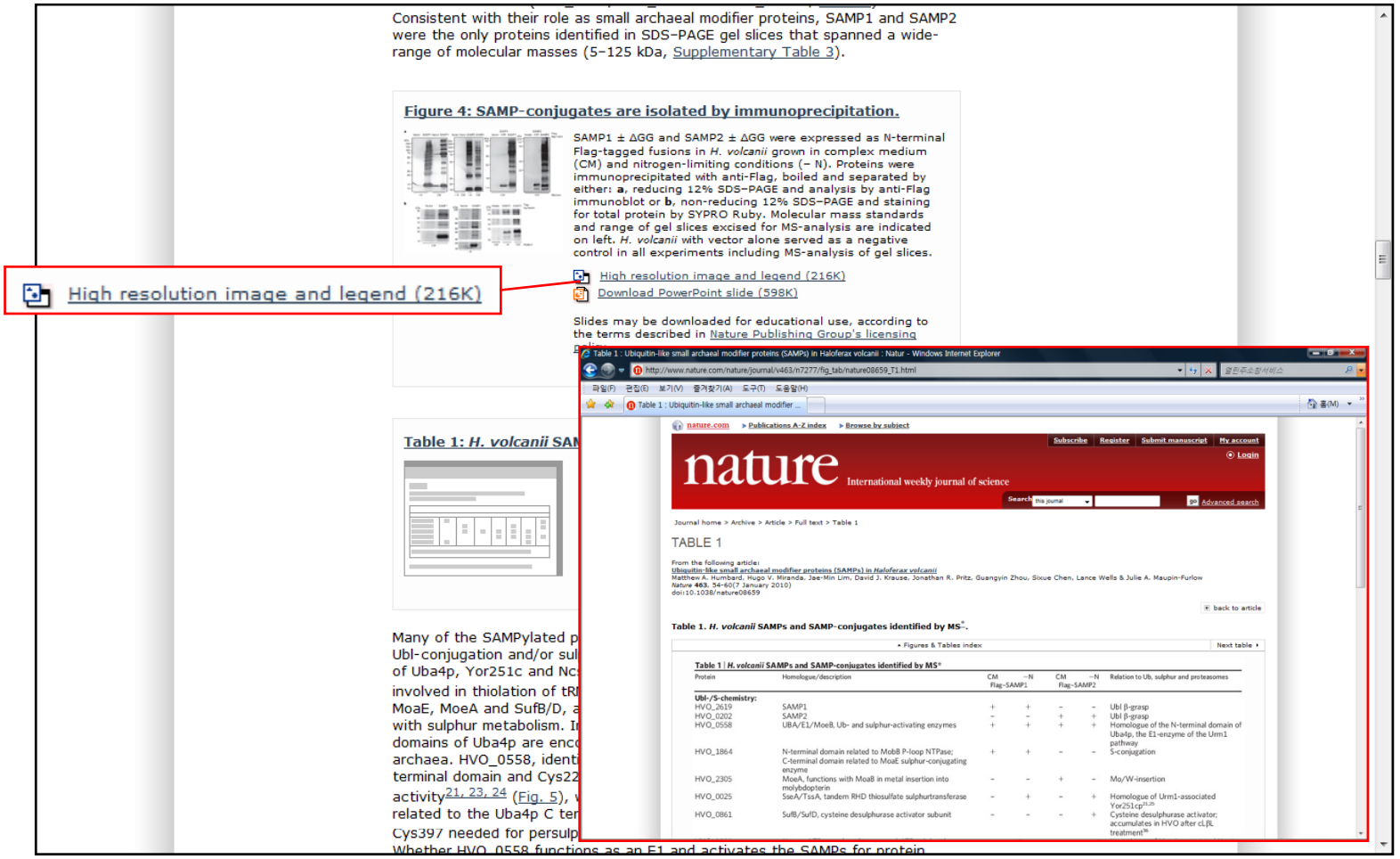

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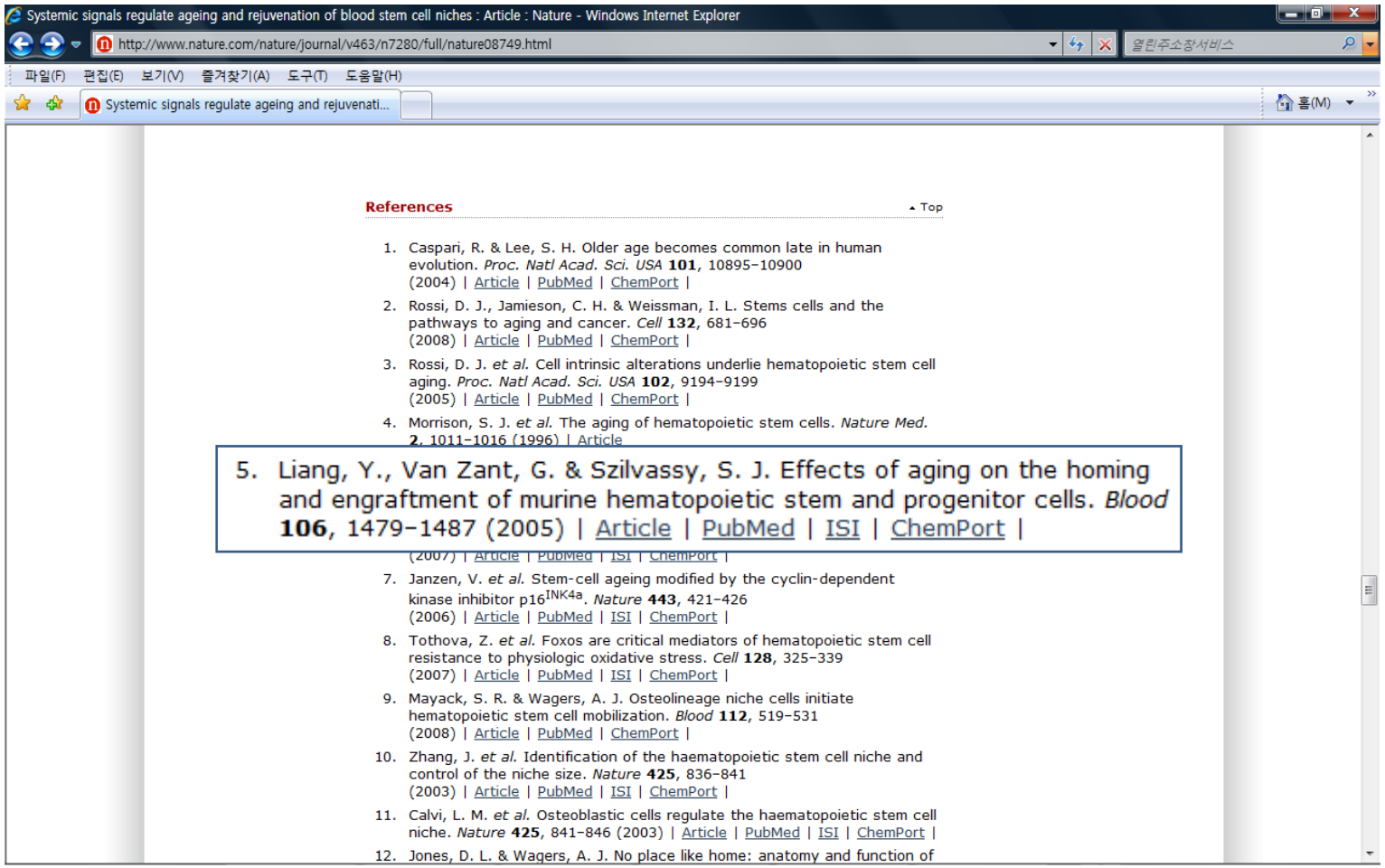

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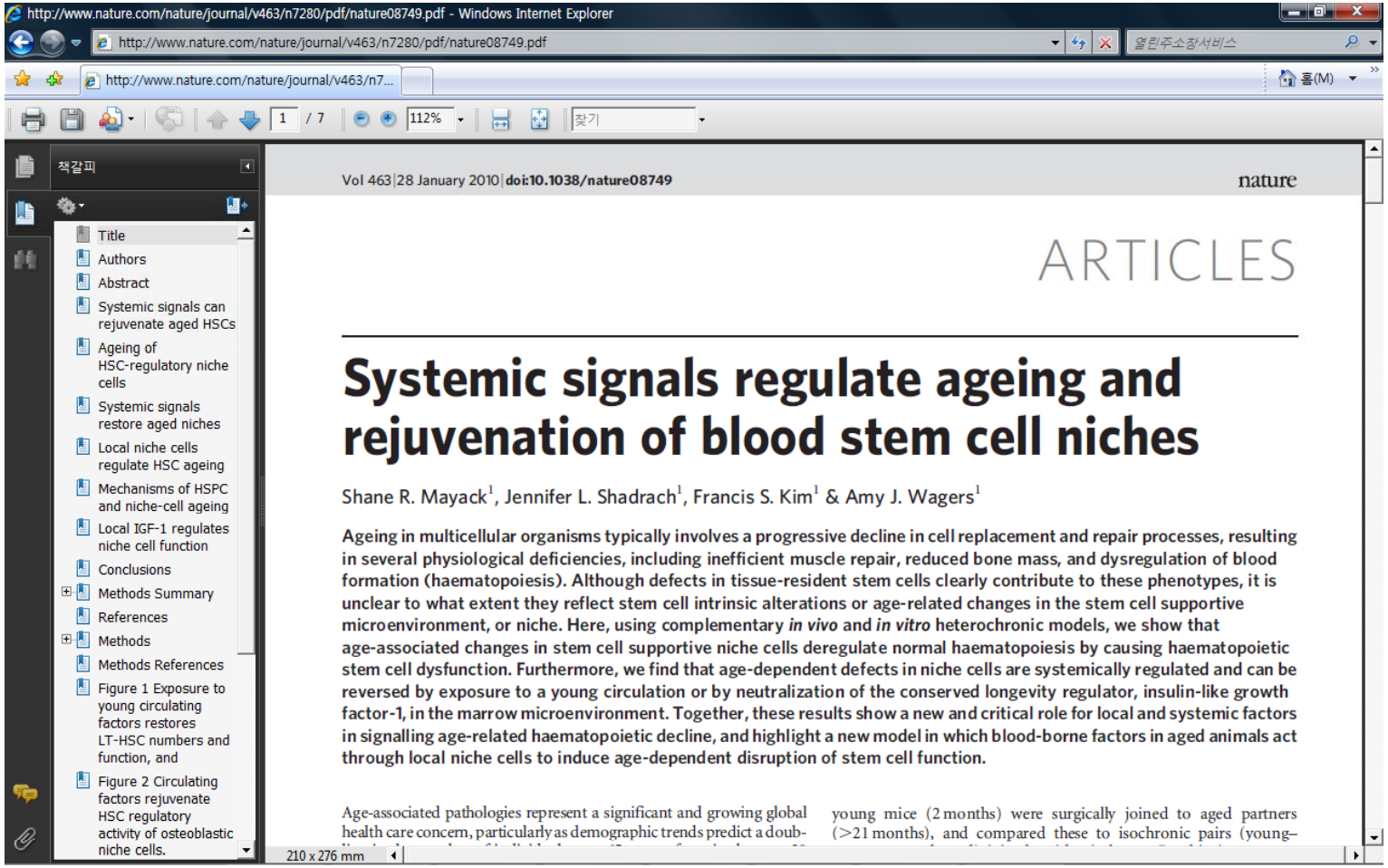
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Abstract
Systemic signals can rejuvenate aged HSCs
Ageing of HSC-regulatory niche cells
Systemic signals restore aged niches
Local niche cells regulate HSC ageing
Mechanisms of HSPC and niche-cell ageing
Local IGF-1 regulates niche cell function
Conclusions
Methods Summary
References
Methods
Methods References
Figure 1 Exposure to young circulating factors restores LT-HSC numbers and function, and
Figure 2 Circulating factors rejuvenate HSC regulatory activity of osteoblastic niche cells.
Vol 463|28 January 2010|doi:10.1038/nature08749
nature
ARTICLES
Systemic signals regulate ageing and rejuvenation of blood stem cell niches
Shane R. Mayack[1], Jennifer L. Shadrach[1], Francis S. Kim[1] & Amy J. Wagers[1]
Ageing in multicellular organisms typically involves a progressive decline in cell replacement and repair processes, resulting in several physiological deficiencies, including inefficient muscle repair, reduced bone mass, and dysregulation of blood formation (haematopoiesis). Although defects in tissue-resident stem cells clearly contribute to these phenotypes, it is unclear to what extent they reflect stem cell intrinsic alterations or age-related changes in the stem cell supportive microenvironment, or niche. Here, using complementary in vivo and in vitro heterochronic models, we show that age-associated changes in stem cell supportive niche cells deregulate normal haematopoiesis by causing haematopoietic stem cell dysfunction. Furthermore, we find that age-dependent defects in niche cells are systemically regulated and can be reversed by exposure to a young circulation or by neutralization of the conserved longevity regulator, insulin-like growth factor-1, in the marrow microenvironment. Together, these results show a new and critical role for local and systemic factors in signalling age-related haematopoietic decline, and highlight a new model in which blood-borne factors in aged animals act through local niche cells to induce age-dependent disruption of stem cell function.
Age-associated pathologies represent a significant and growing global health care concern, particularly as demographic trends predict a doub-
young mice (2 months) were surgically joined to aged partners (>21 months), and compared these to isochronic pairs (young–
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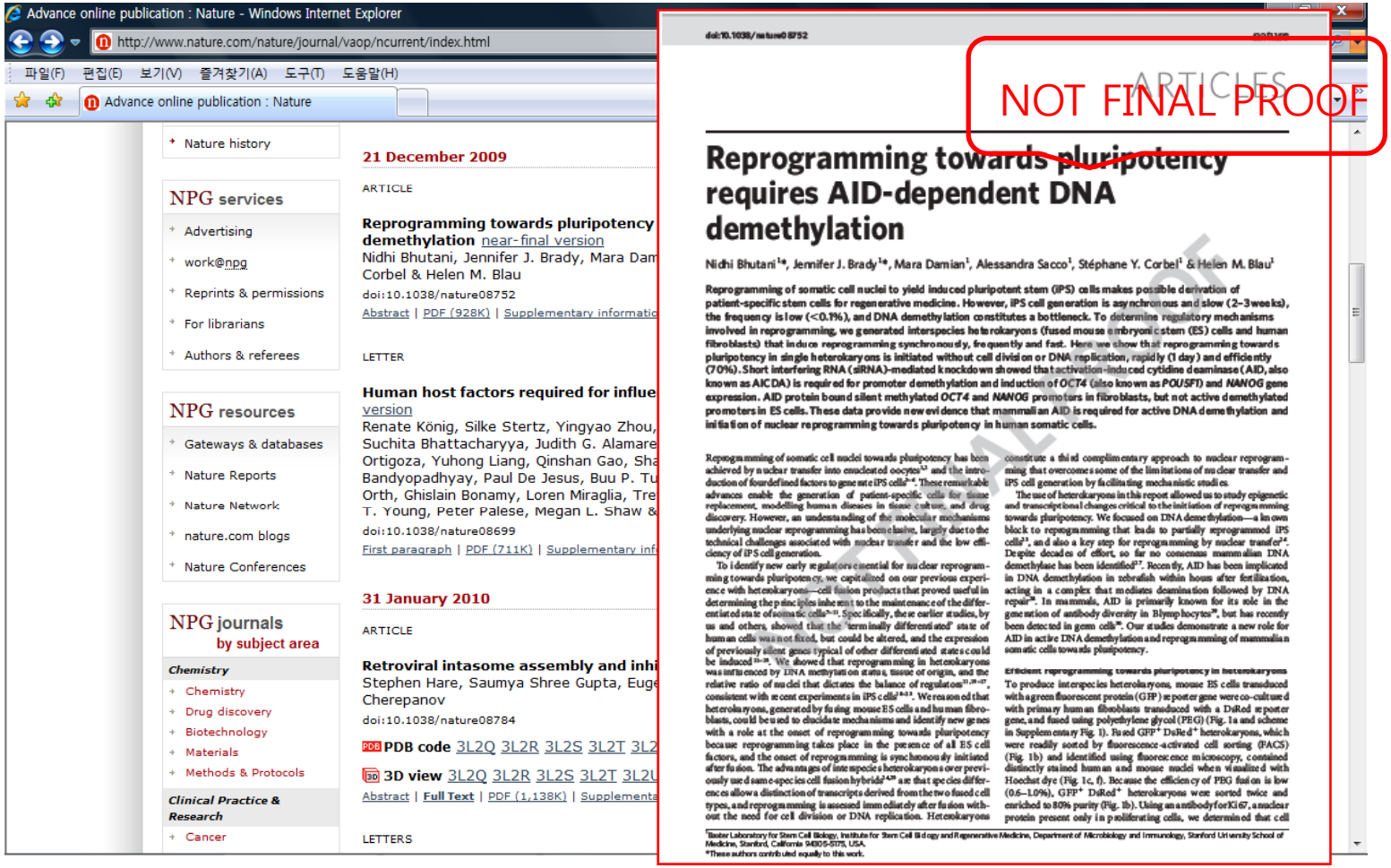

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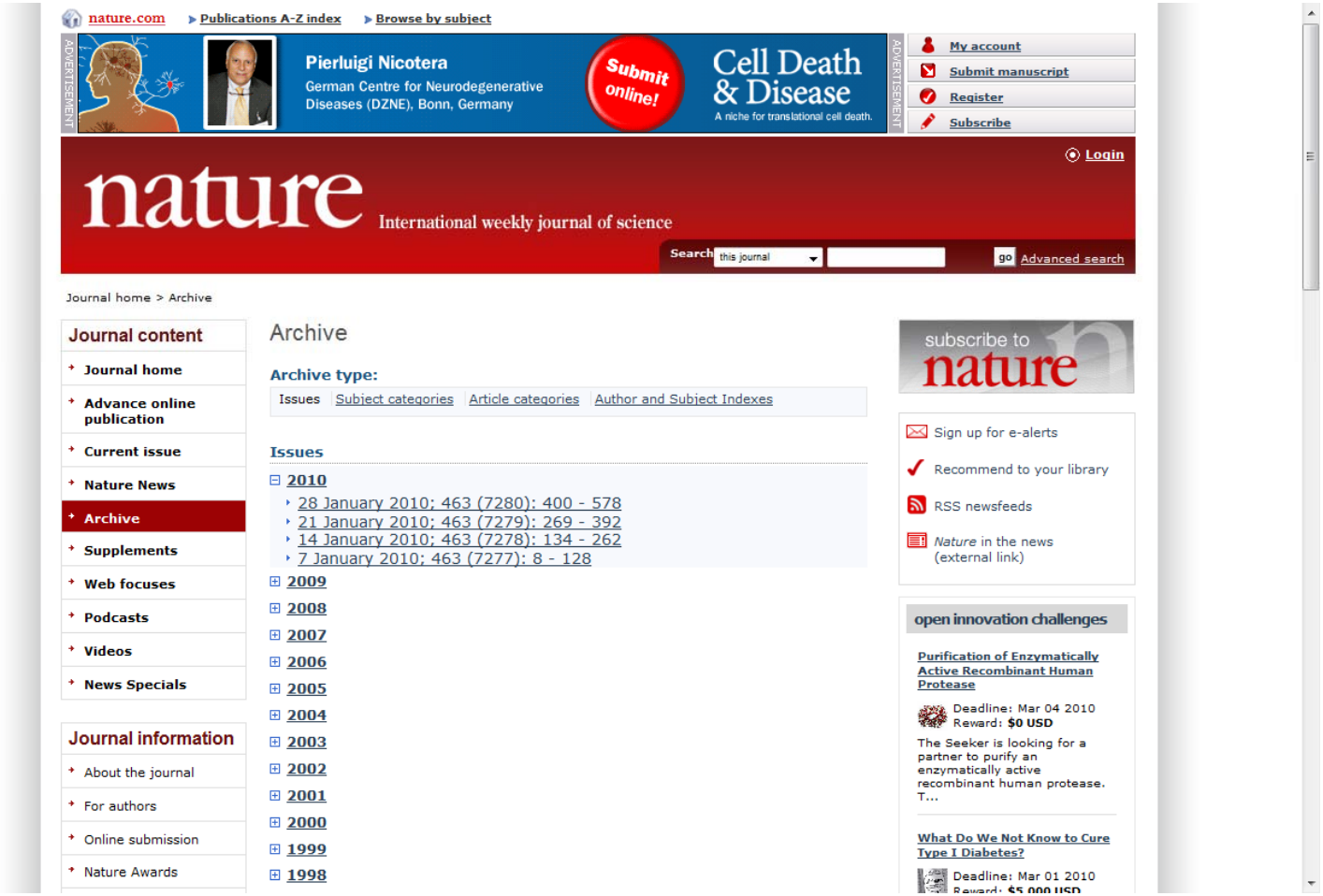

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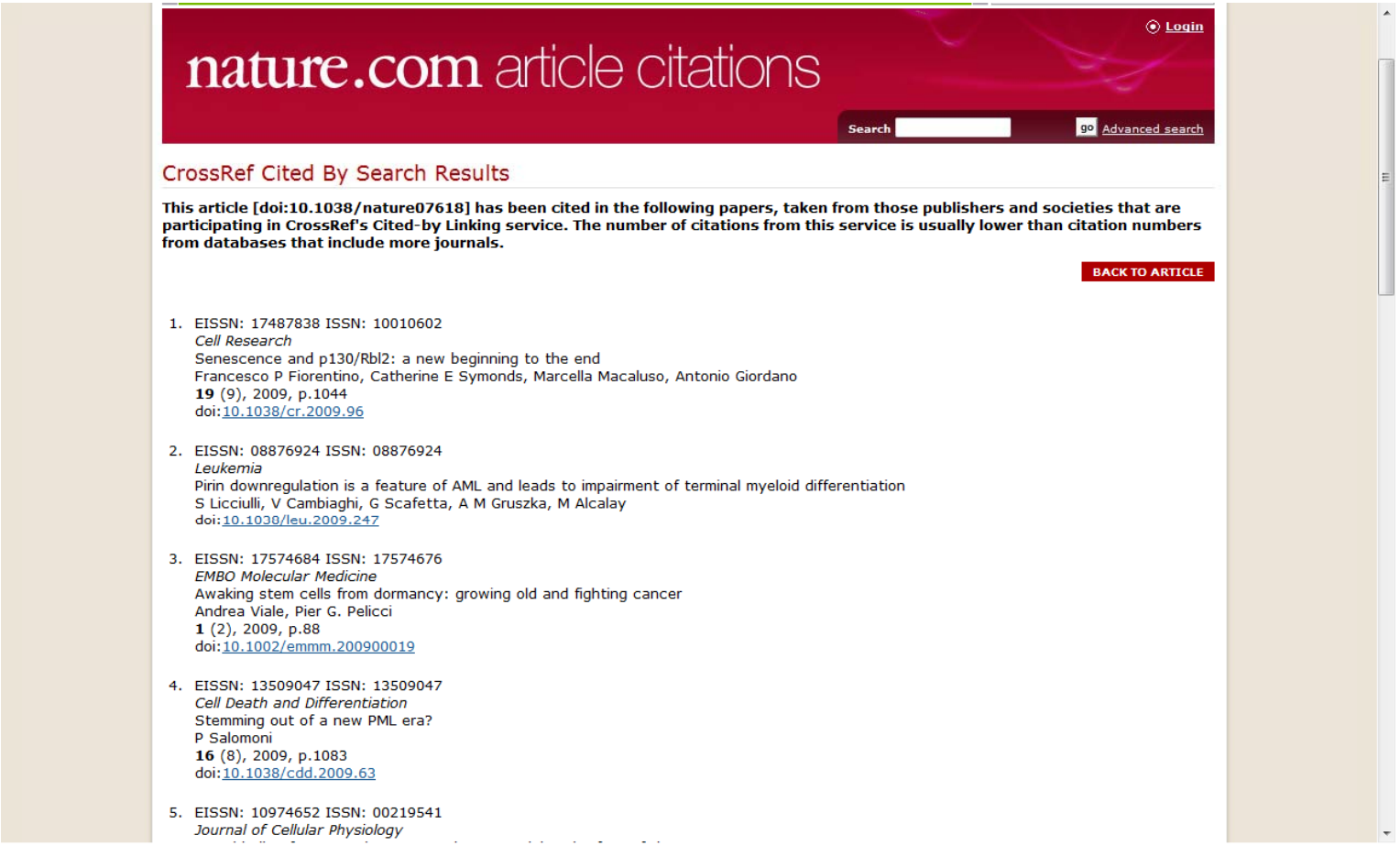

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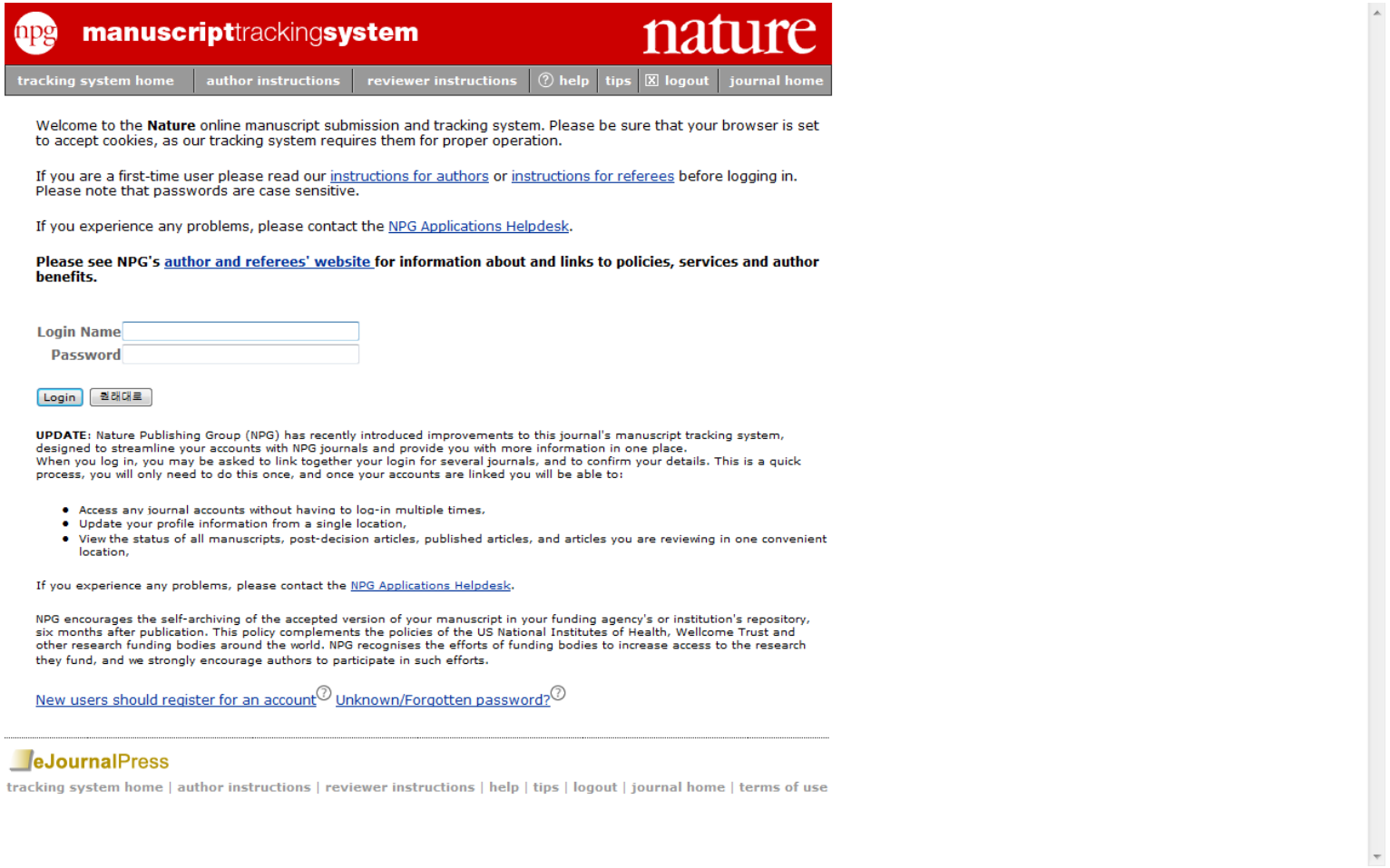

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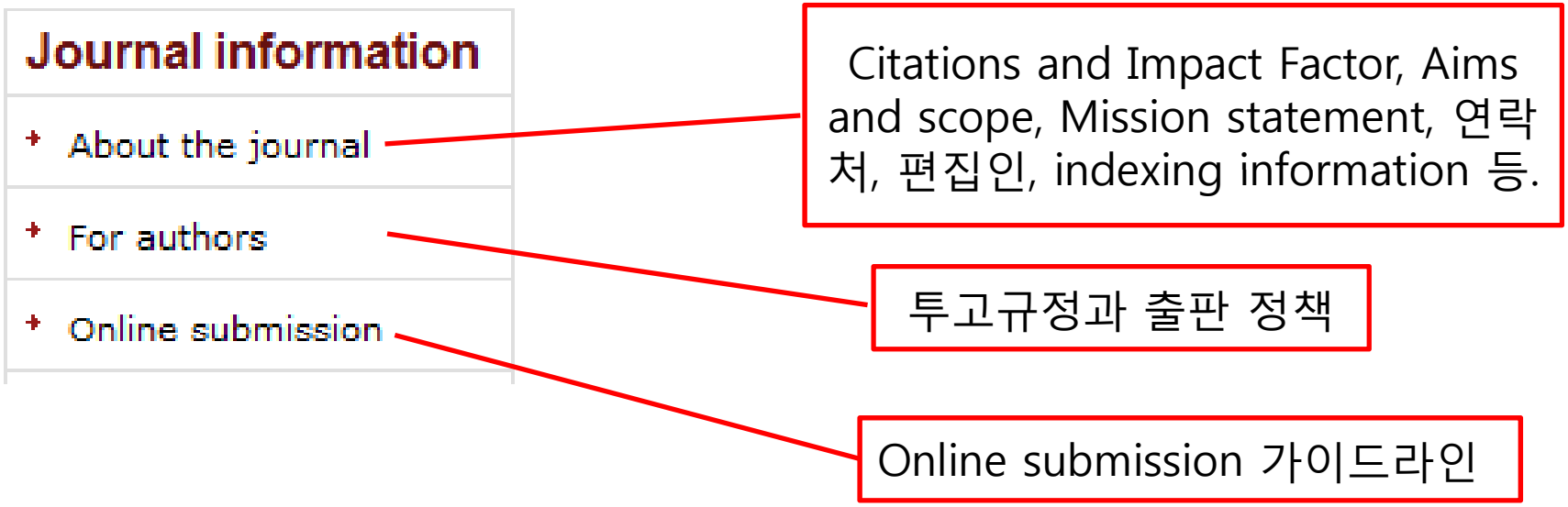

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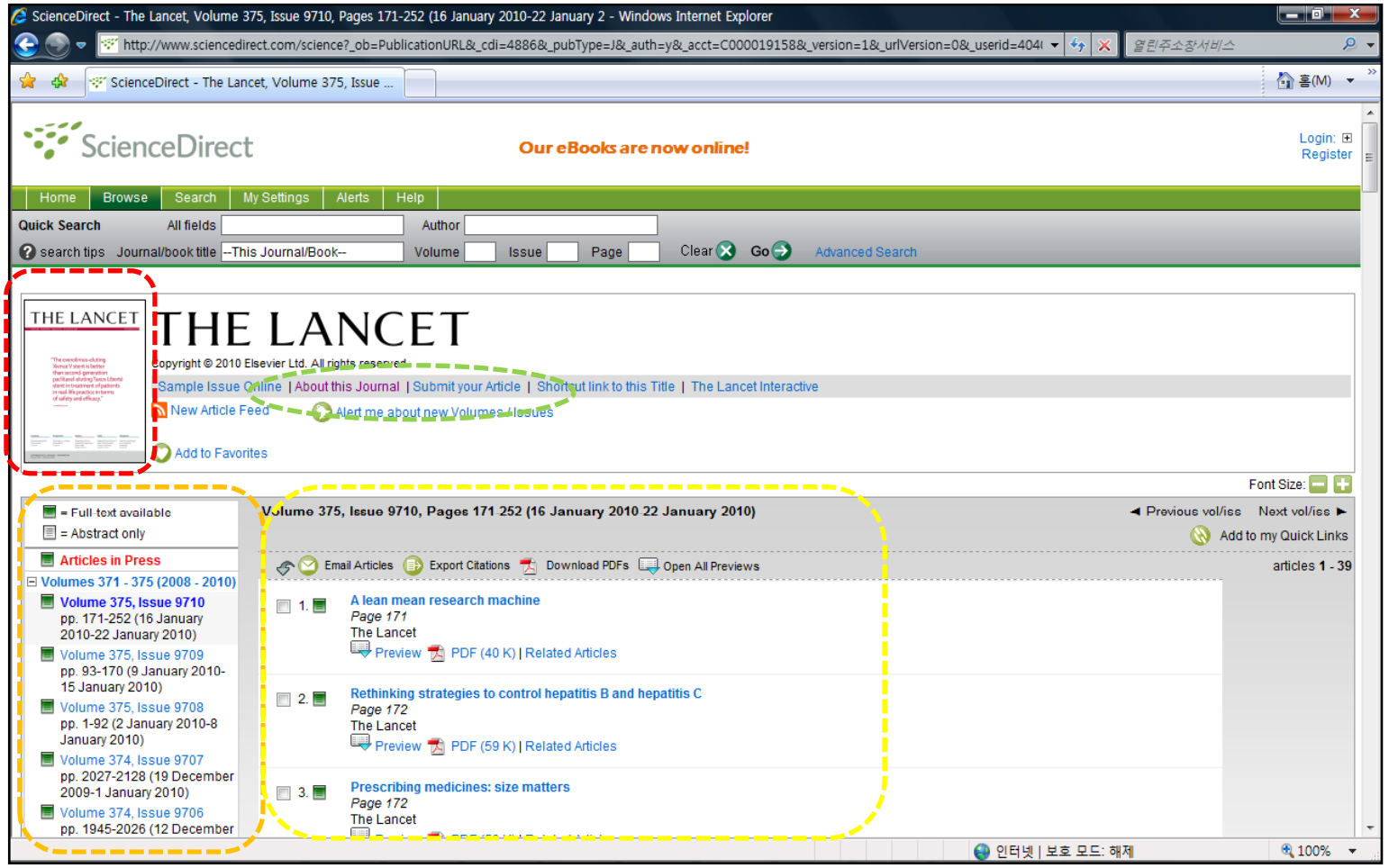

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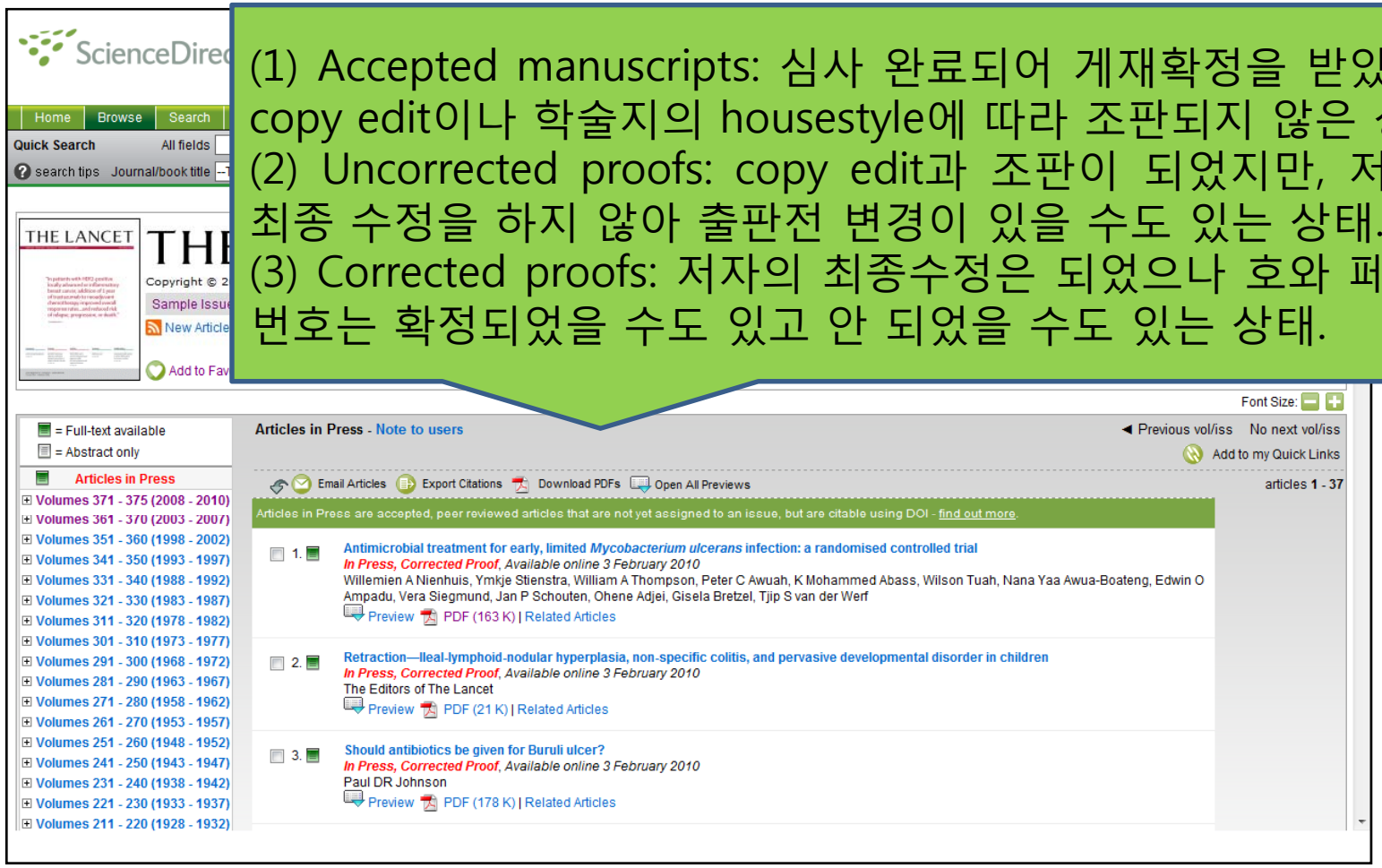

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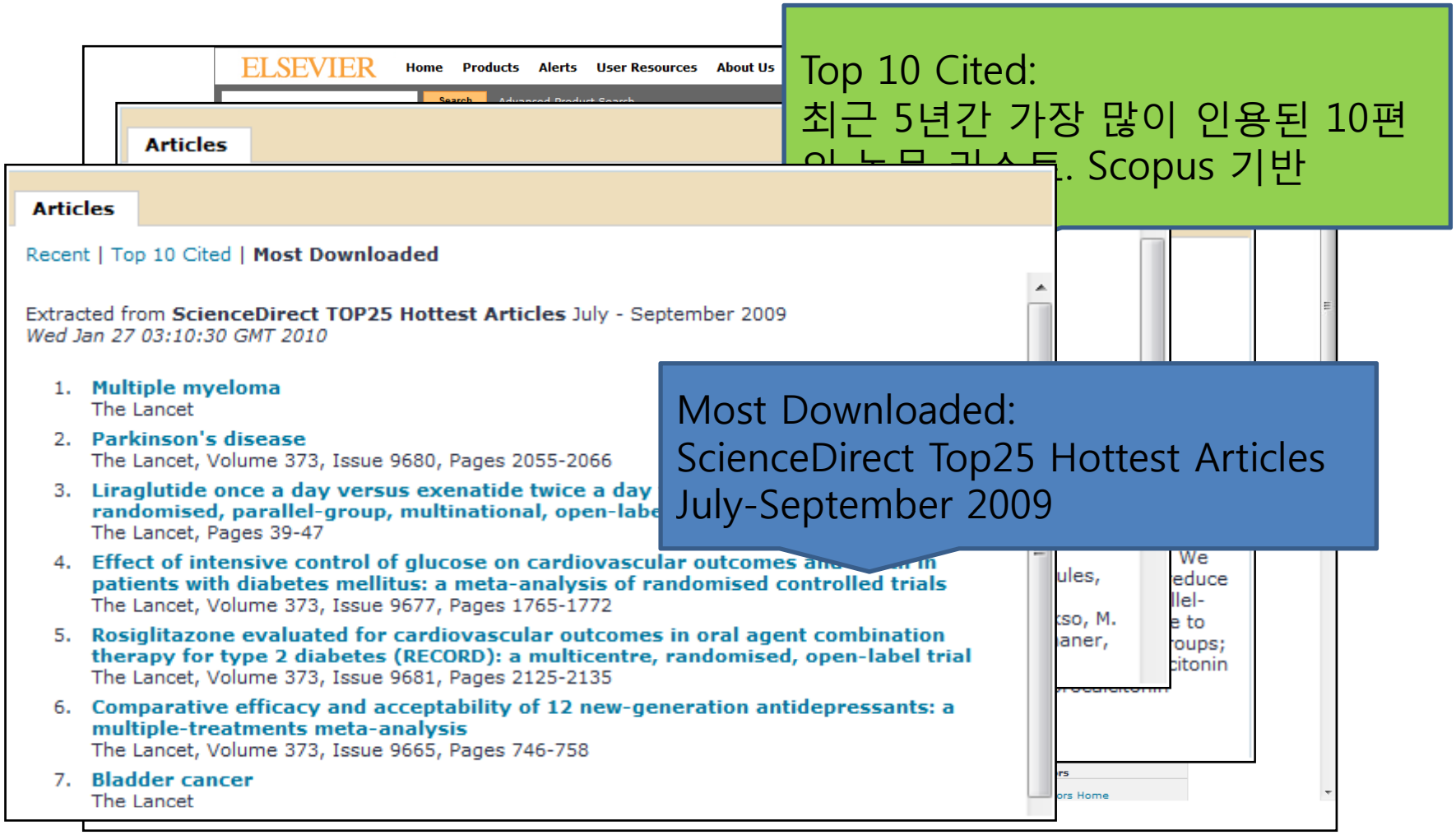

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group, 1 in exenatide group), and they were included in the safety but not intention-to-treat populations. 33 of 235 participants withdrew from liraglutide and 45 of 232 from exenatide treatment; withdrawal rates were not significantly different between groups. Adverse events were the most common reason for withdrawal in both groups. The characteristics of the study population were typical for participants with type 2 diabetes, and baseline characteristics were well matched between treatment groups (table 1).

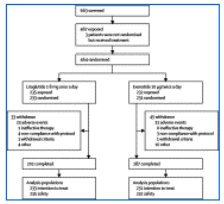

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Figure 1. Trial profile

Of the adverse events leading to withdrawal, nausea was the most common (14 patients in the liraglutide group and 16 in the exenatide group). Participants were exposed to treatment if they had received at least one dose of study medication.

Table 1.

Baseline demographic and disease characteristics

	Liraglutide 1·8 mg once a day (n=233)	Exenatide 10 μg twice a day (n=231)
Men	114 (49%)	127 (55%)
Age (years)	56·3 (9·8)	57·1 (10·8)
Race		
White	216 (93%)	210 (91%)
Asian/Pacific Islander	1 (<1%)	5 (2%)
Black*	13 (6%)	12 (5%)
Other	3 (1%)	4 (2%)
Hispanic or Latin American ethnic origin	32 (14%)	25 (11%)
Weight (kg)	93·1 (20·1)	93·0 (19·5)
Body-mass index (kg/m²)	32·9 (5·5)	32·9 (5·7)
Duration of diabetes (years)	8·5 (6·2)	7·9 (5·9)

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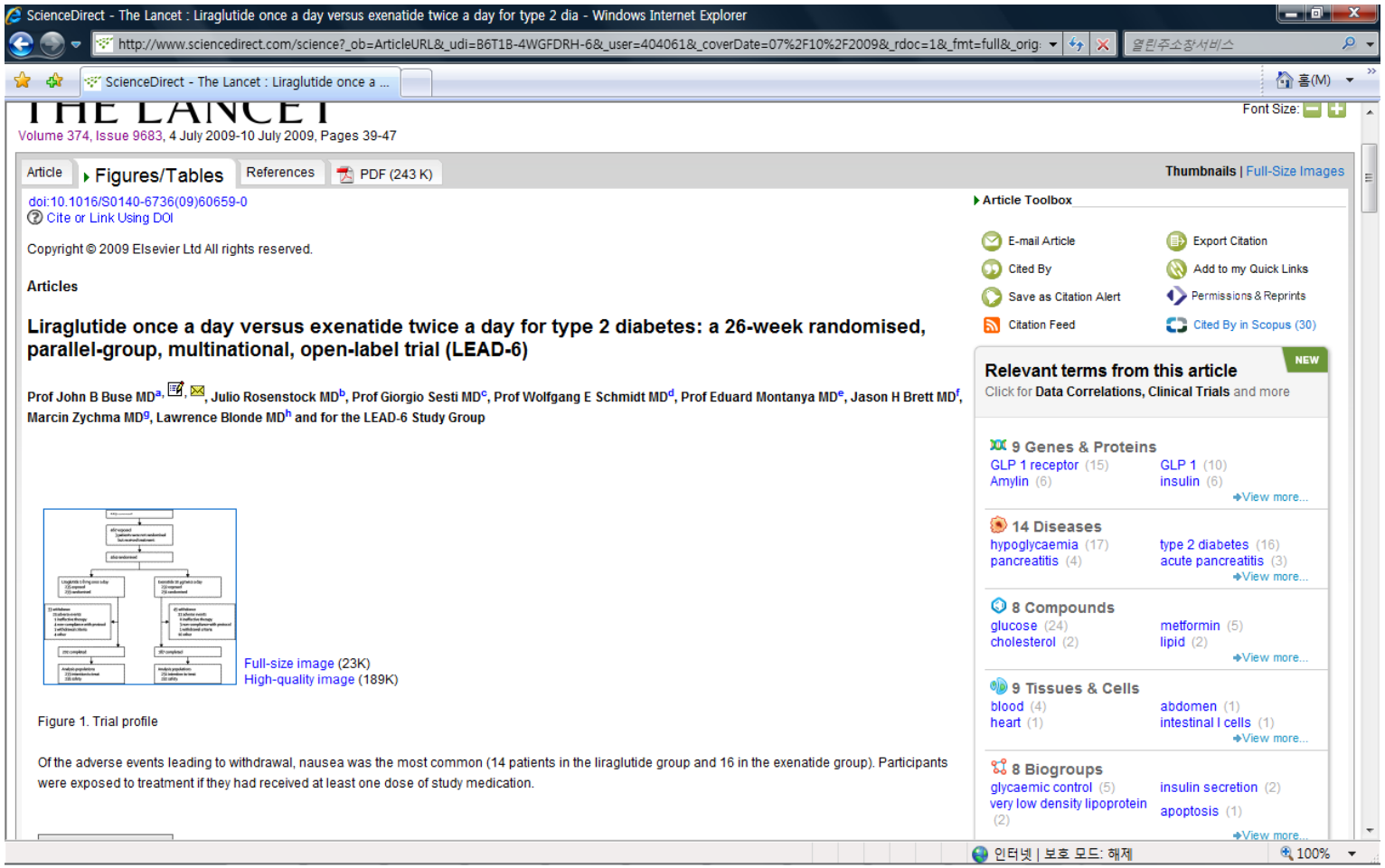

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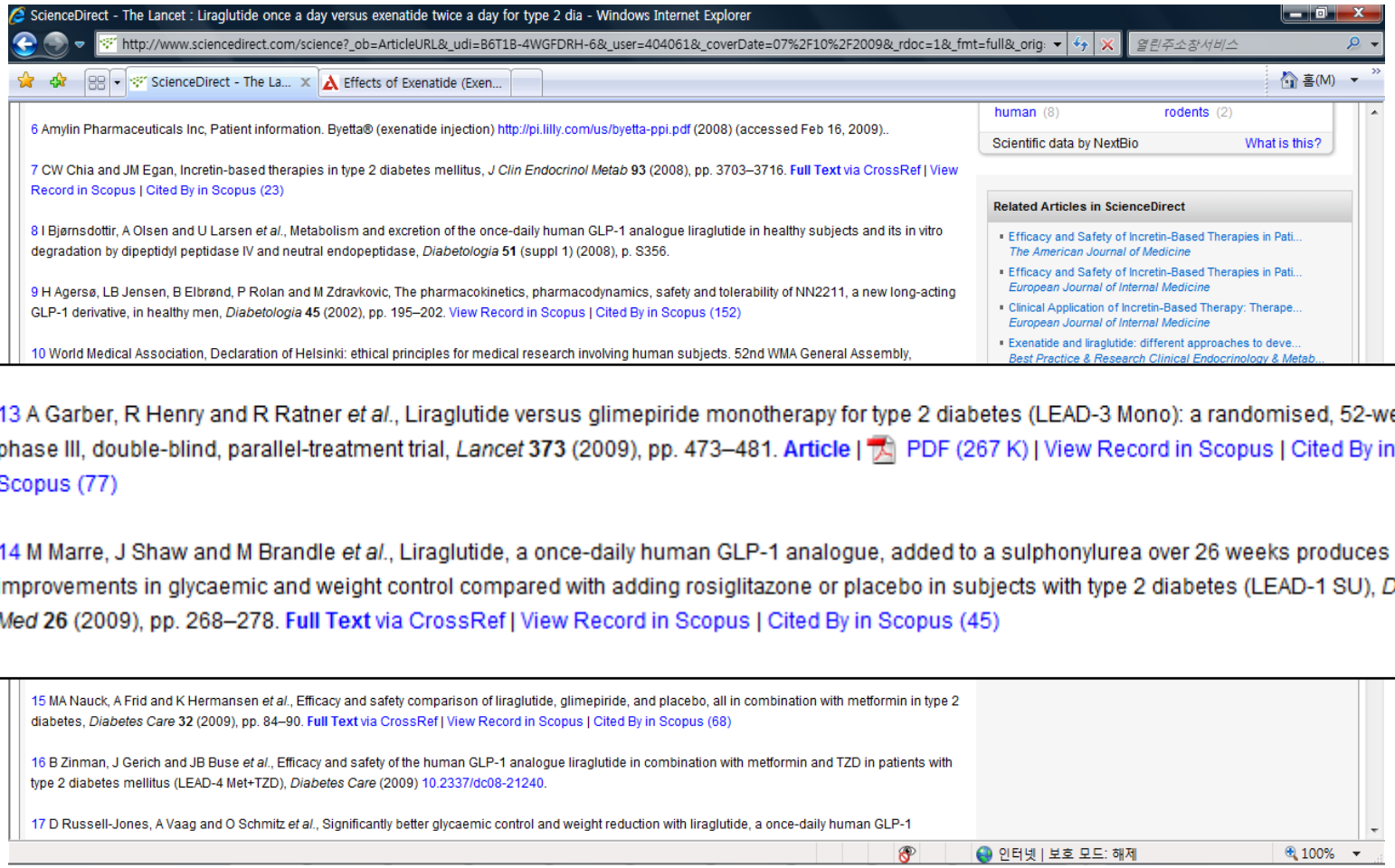

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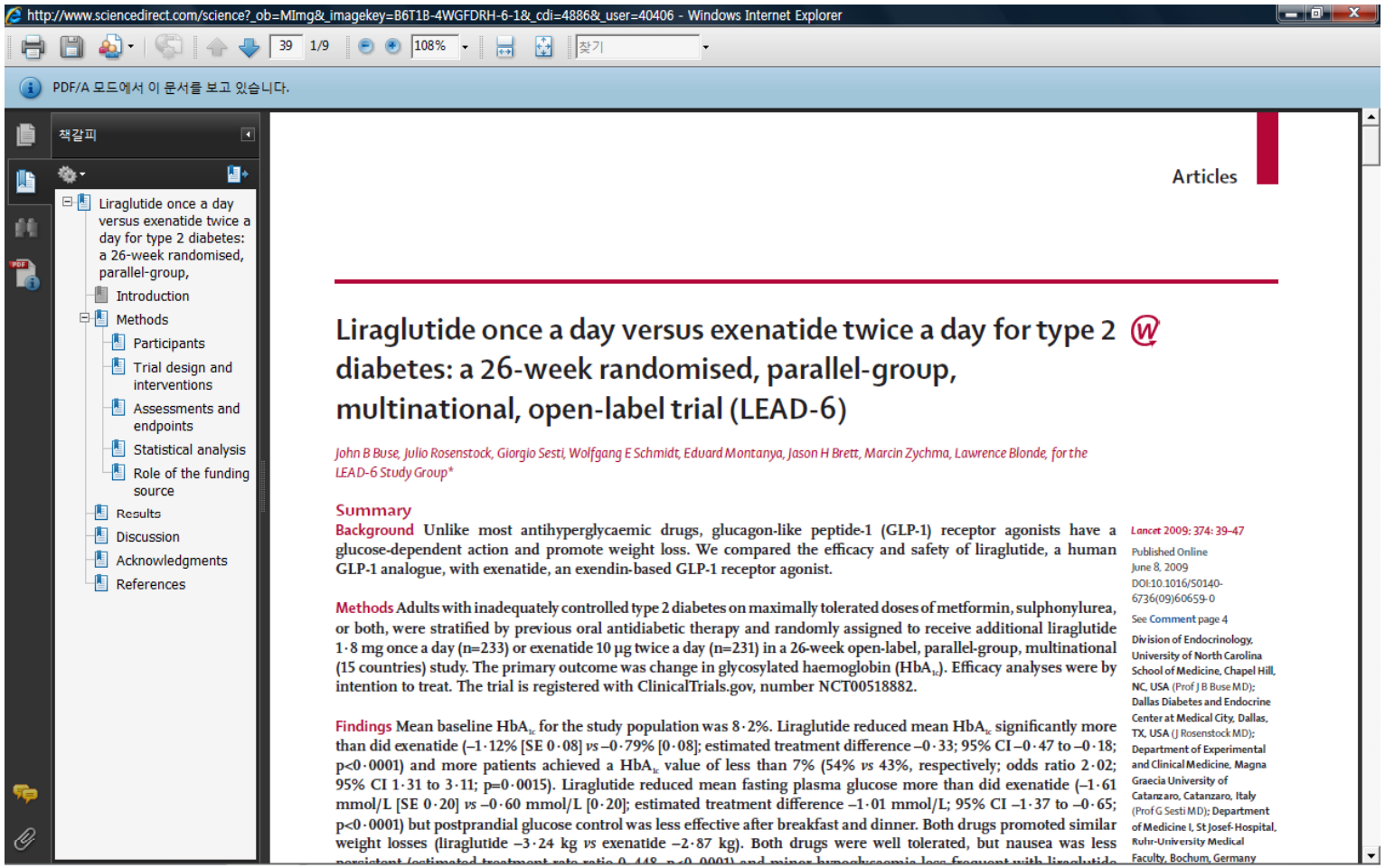

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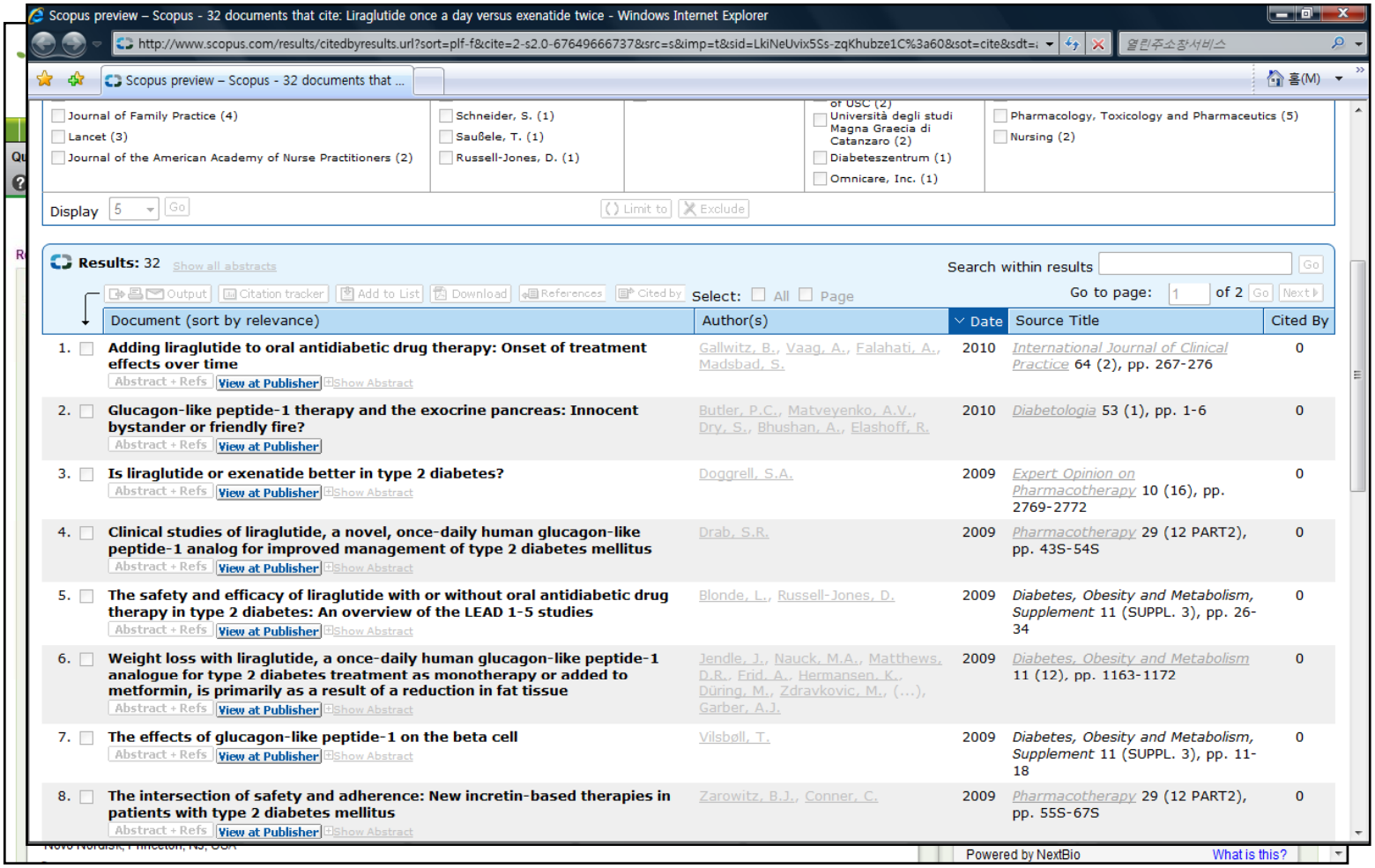

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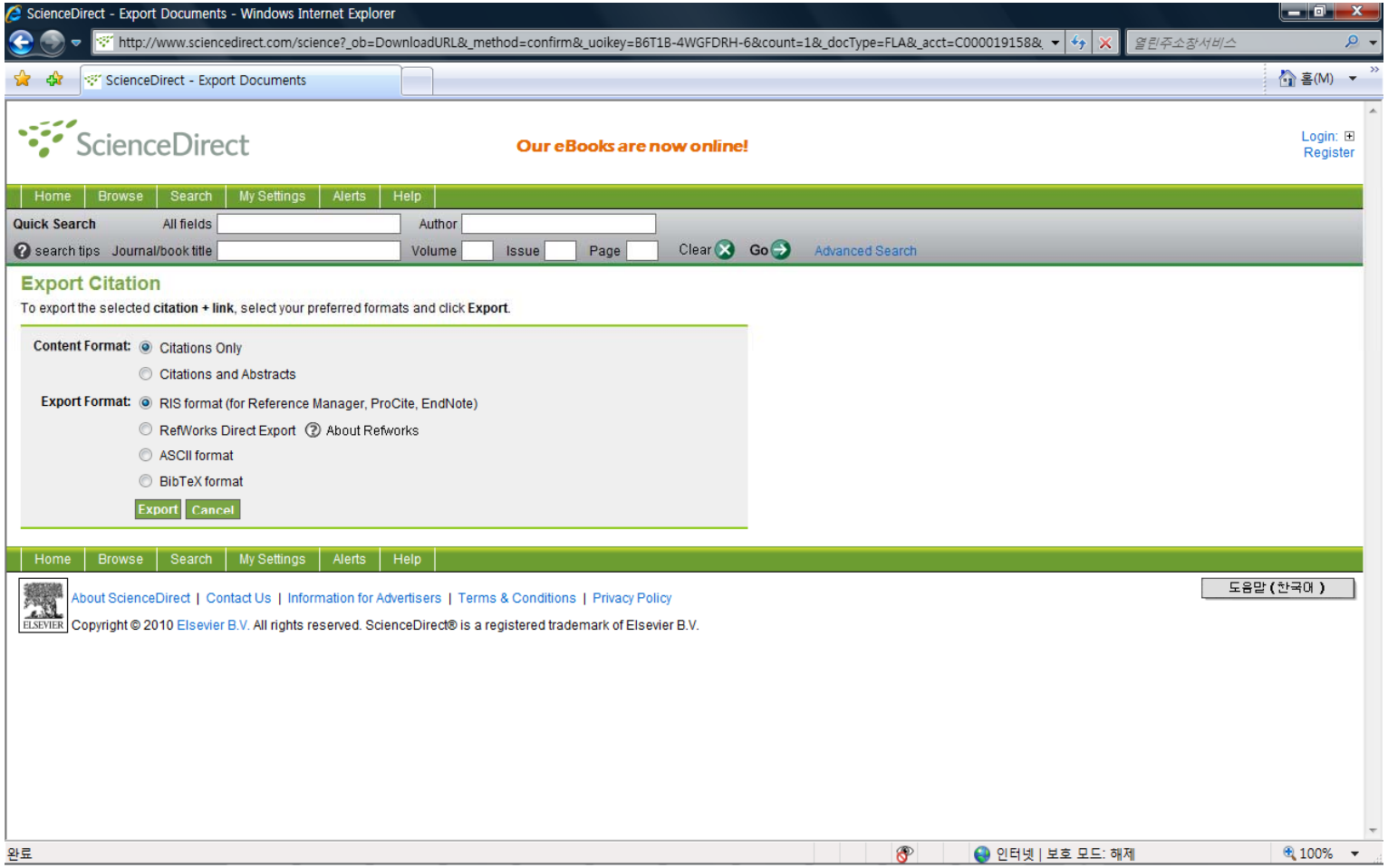

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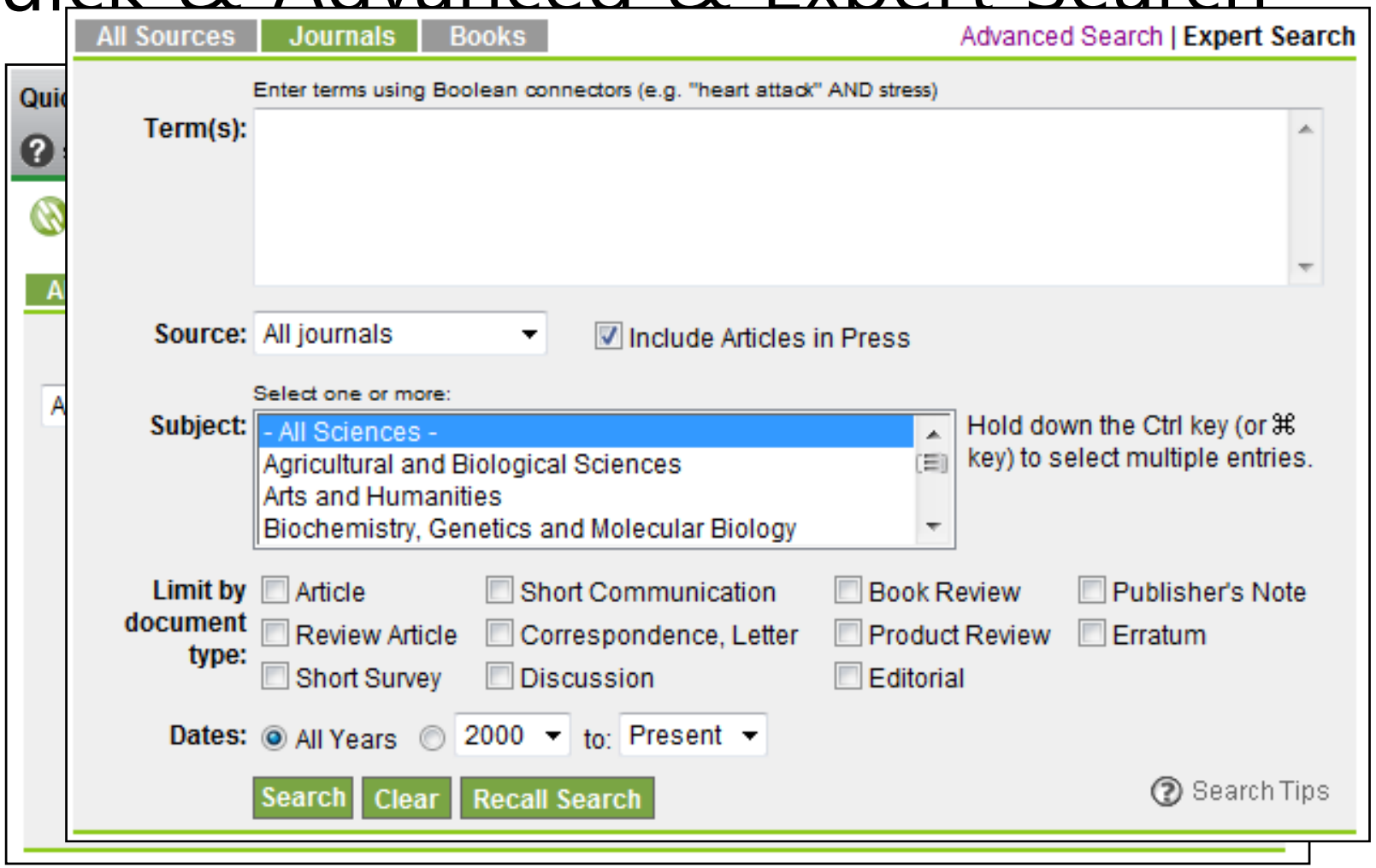

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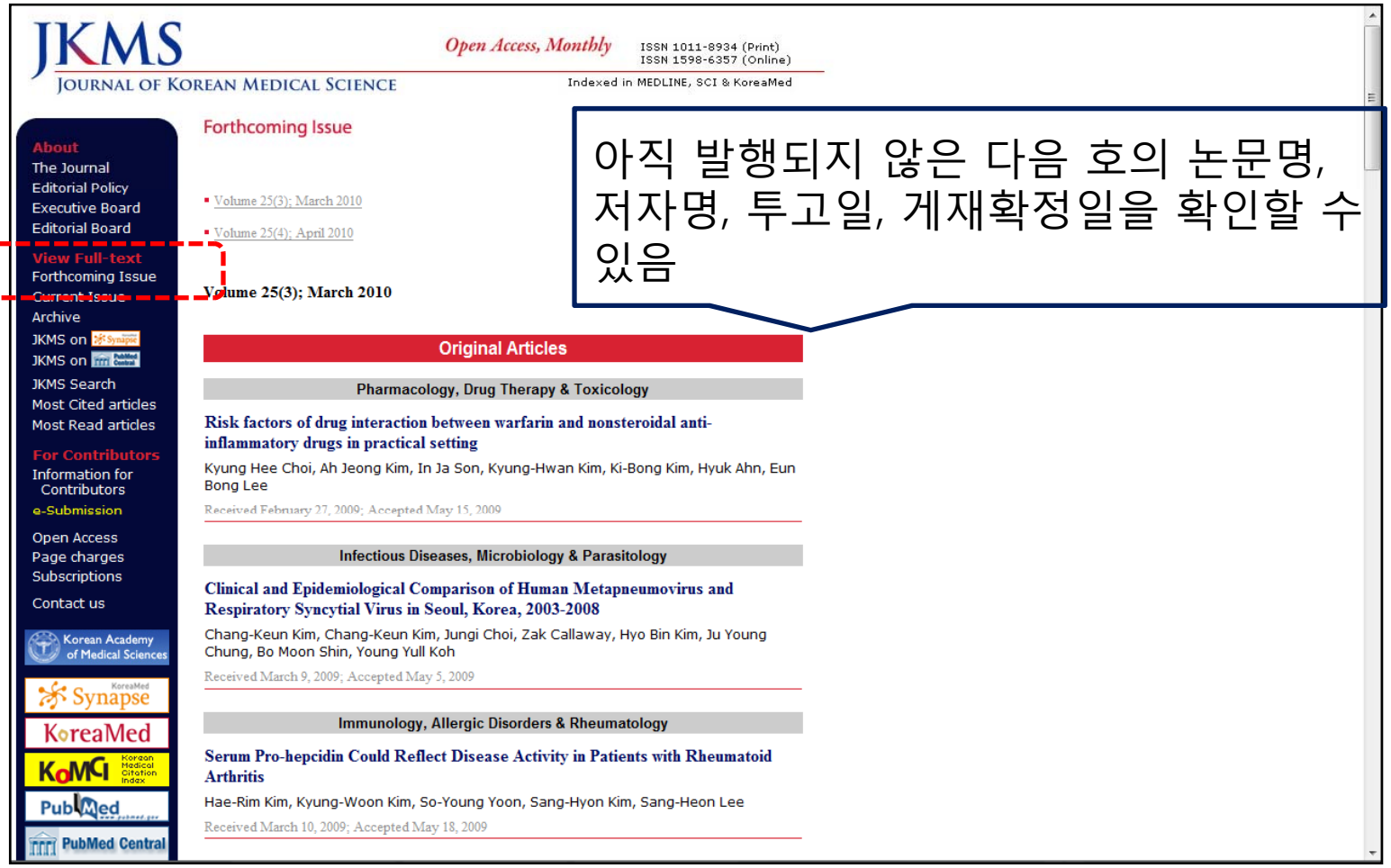

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Review

Update on the management of antibiotic allergy

Bernard Yu-Hor Thong.

Department of Rheumatology, Allergy and Immunology, Tar

Received October 18, 2009; Accepted October 19, 2009.

Abstract posted online 2010 January 5

Original Articles

Clinical characteristics according to sensitized allerg bronchial asthma

Jae Woo Jung, Jae Chol Choi, Jong Wook Shin, Jae Yeo

Department of Internal Medicine, Chung-Ang University Co

Received August 5, 2009; Accepted December 16, 2009.

Abstract posted online 2010 January 22

IL-13 gene polymorphisms are associated with rhino inflammation in aspirin intolerant asthma

Nami Shrestha Palikhe, Seung-Hyun Kim, Bo-Young Cho, Gi Hae-Sim Park.

Department of Allergy and Rheumatology, Ajou University

Received August 31, 2009; Accepted November 26, 2009.

Abstract posted online 2010 January 5

Allergy Asthma Immunol Res. 2010 Forthcoming.
Abstract posted online 2010 January 5.

Update on the management of antibiotic allergy

Bernard Yu-Hor Thong

Department of Rheumatology, Allergy and Immunology, Tan Tock Seng Hospital, Singapore.

Received October 18, 2009; Accepted October 19, 2009.

Abstract

Drug allergy to antibiotics may occur in the form of immediate or non-immediate (delayed) hypersensitivity reactions. Immediate reactions are usually IgE-mediated whereas non-immediate hypersensitivity reactions are usually non-IgE or T-cell mediated. The clinical manifestations of antibiotic allergy may be cutaneous, organ-specific (e.g. blood dyscracias, hepatitis, interstitial nephritis), systemic (e.g. anaphylaxis, drug induced hypersensitivity syndrome) or various combinations of these. Severe cutaneous adverse reactions manifesting as Stevens Johnson syndrome or toxic epidermal necrolysis (TEN) may be potentially life-threatening. The management of antibiotic allergy begins with the identification of the putative antibiotic from a detailed and accurate drug history, complemented by validated in-vivo and in-vitro allergological tests. This will facilitate avoidance of the putative antibiotic through patient education, use of drug alert cards, and electronic medical records with in-built drug allergy/adverse drug reaction prescription and dispensing checks. Knowledge of the evidence for specific antibiotic cross-reactivities is also important in patient education. Apart from withdrawal of the putative antibiotic, immunomodulatory agents like high-dose intravenous immunoglobulins may have a role in TEN. Drug desensitization where the benefits outweigh the risks, and where no alternative antibiotics can be used for various reasons, may be considered in certain situations. Allergological issues pertaining to electronic drug allergy alerts, computerized physician prescriptions and decision support systems, and antibiotic de-escalation in antimicrobial stewardship programmes are also discussed.

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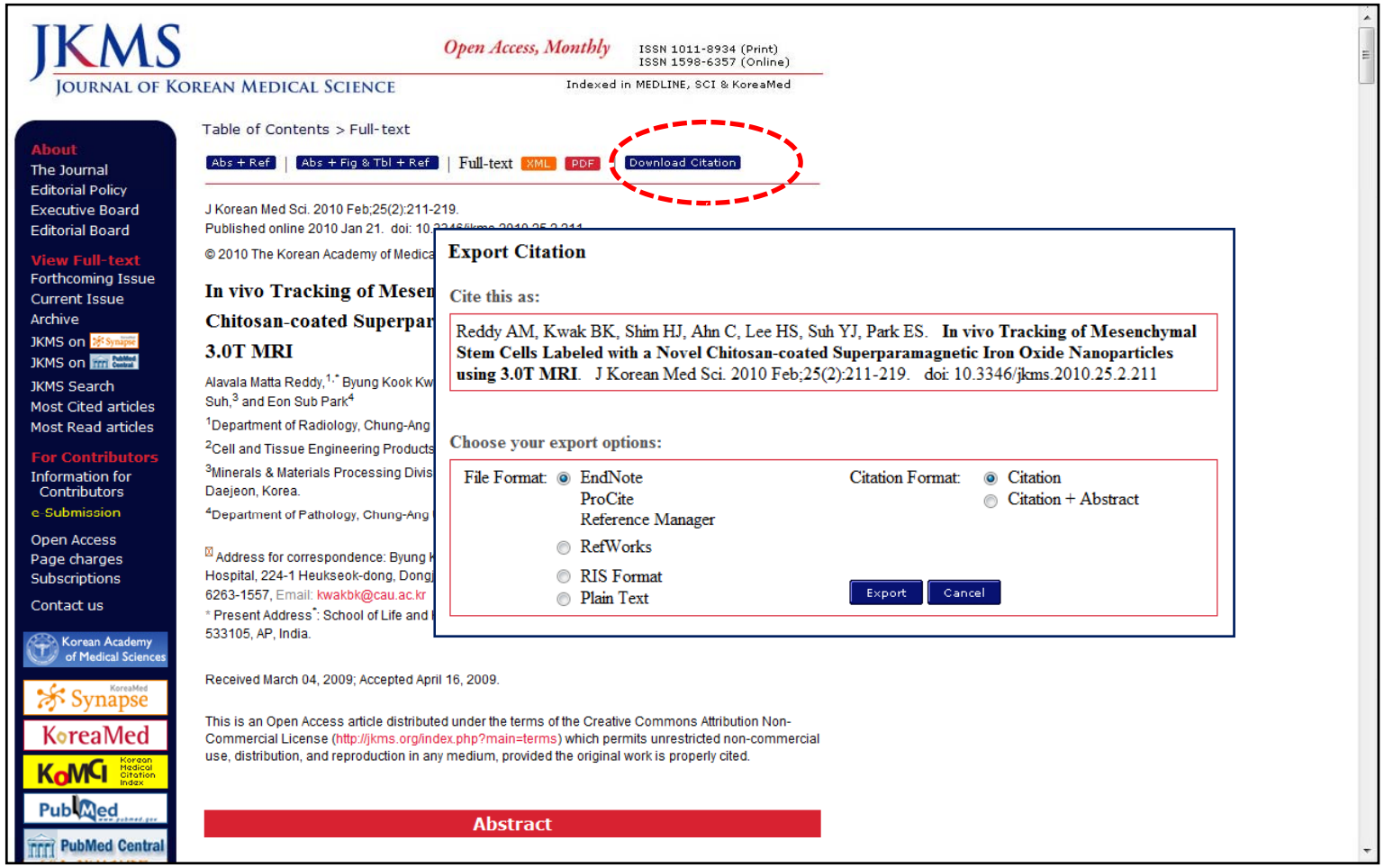

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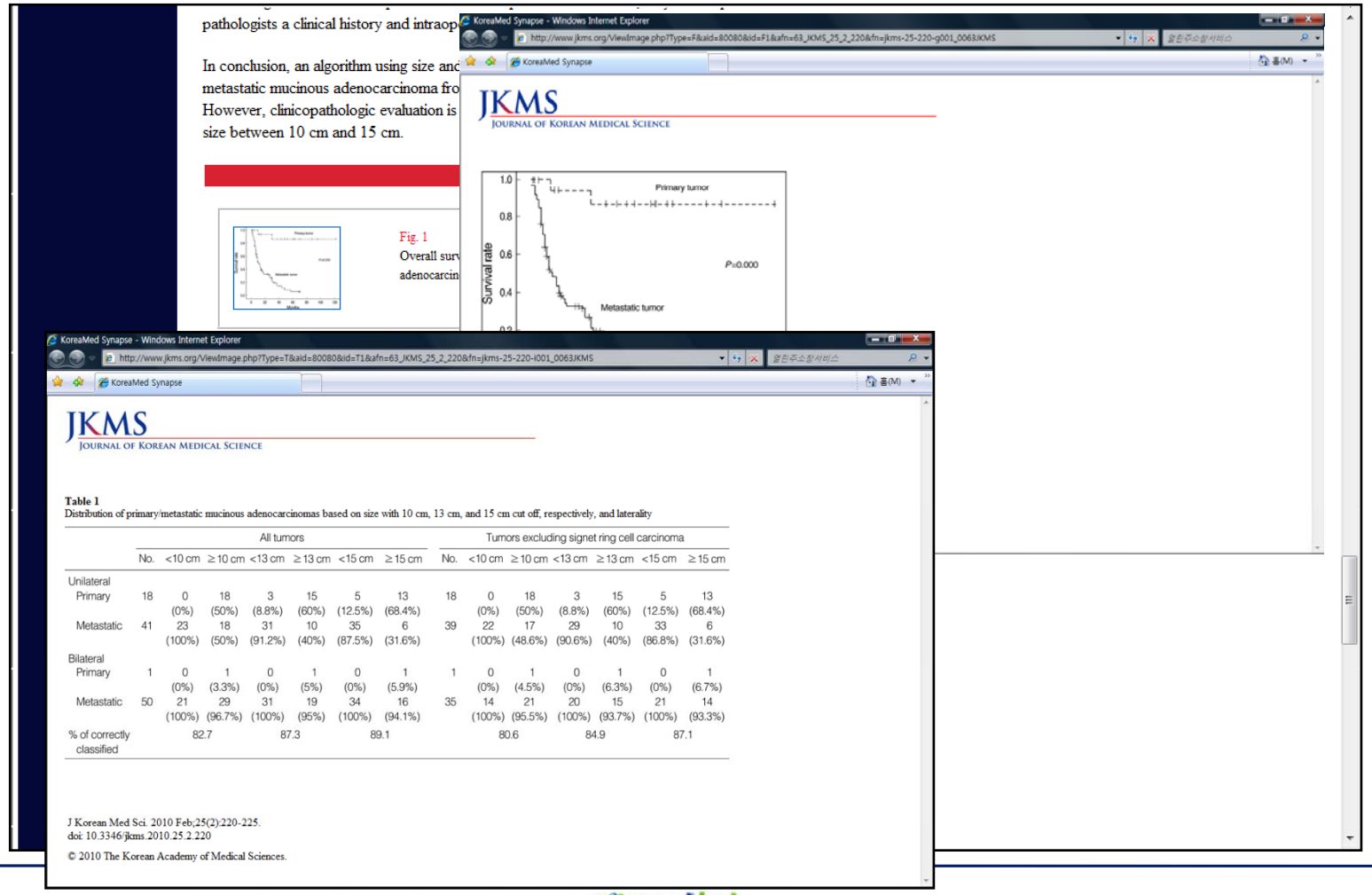

Table 1
Distribution of primary/metastatic mucinous adenocarcinomas based on size with 10 cm, 13 cm, and 15 cm cut off, respectively, and laterality

	All tumors							Tumors excluding signet ring cell carcinoma						
	No.	<10 cm	≥10 cm	<13 cm	≥13 cm	<15 cm	≥15 cm	No.	<10 cm	≥10 cm	<13 cm	≥13 cm	<15 cm	≥15 cm
Unilateral														
Primary	18	0 (0%)	18 (50%)	3 (8.8%)	15 (60%)	5 (12.5%)	13 (68.4%)	18	0 (0%)	18 (50%)	3 (8.8%)	15 (60%)	5 (12.5%)	13 (68.4%)
Metastatic	41	23 (100%)	18 (50%)	31 (91.2%)	10 (40%)	35 (87.5%)	6 (31.6%)	39	22 (100%)	17 (48.6%)	29 (90.6%)	10 (40%)	33 (86.8%)	6 (31.6%)
Bilateral														
Primary	1	0 (0%)	1 (3.3%)	0 (0%)	1 (5%)	0 (0%)	1 (5.9%)	1	0 (0%)	1 (4.5%)	0 (0%)	1 (6.3%)	0 (0%)	1 (6.7%)
Metastatic	50	21 (100%)	29 (96.7%)	31 (100%)	19 (95%)	34 (100%)	16 (94.1%)	35	14 (100%)	21 (95.5%)	20 (100%)	15 (93.7%)	21 (100%)	14 (93.3%)
% of correctly classified			82.7		87.3		89.1			80.6		84.9		87.1

J Korean Med Sci. 2010 Feb;25(2):220-225.
doi: 10.3346/jkms.2010.25.2.220

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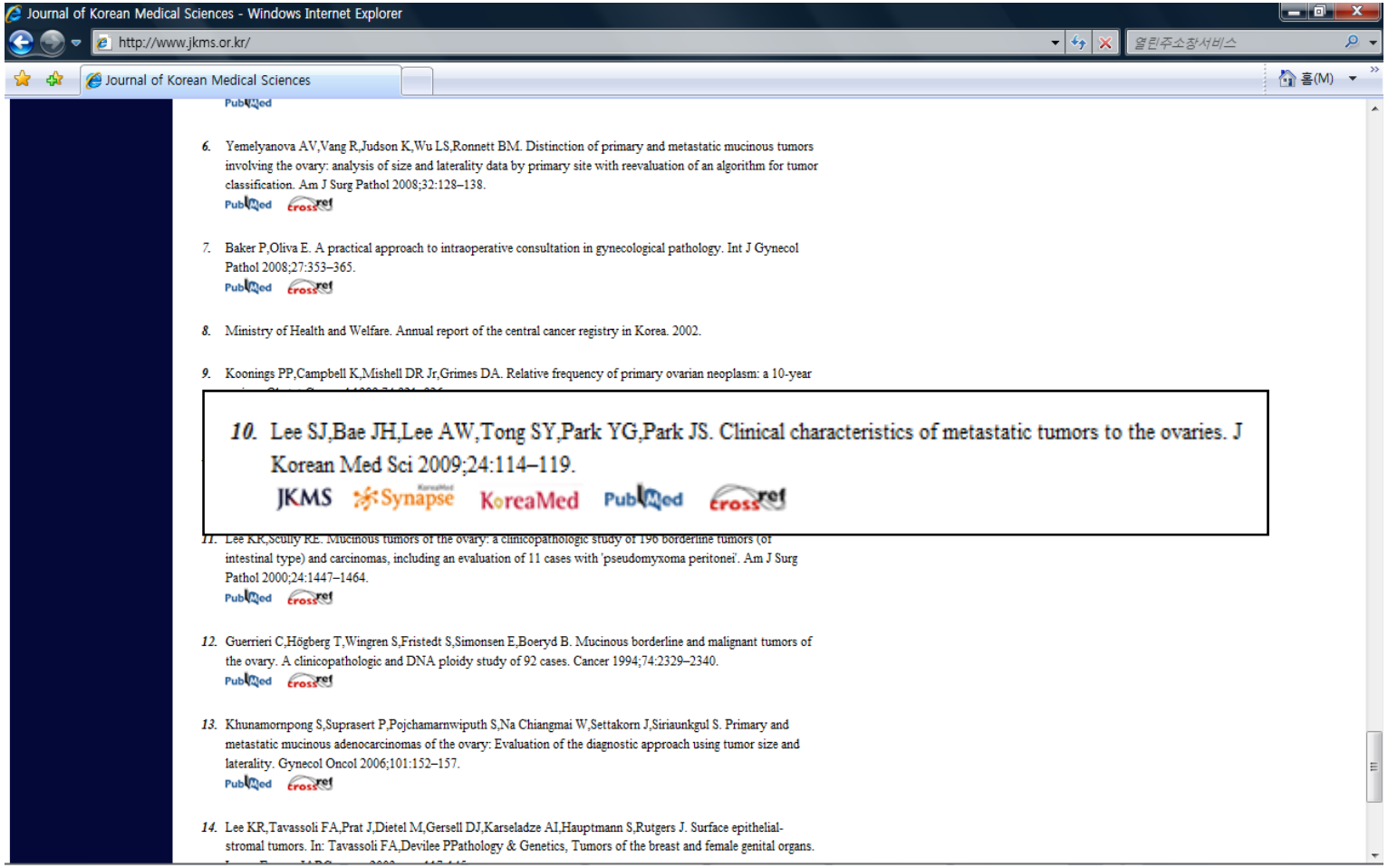

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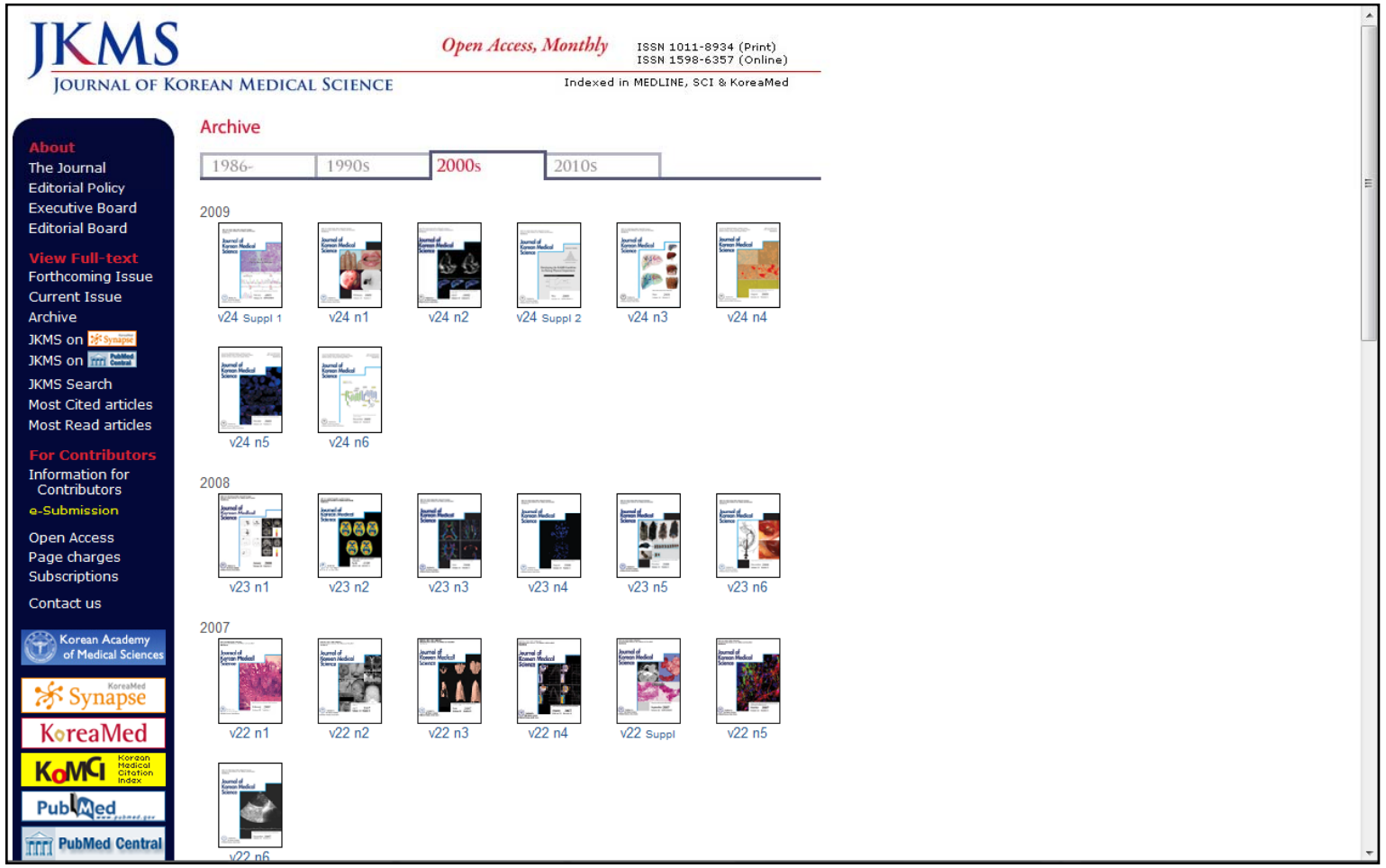

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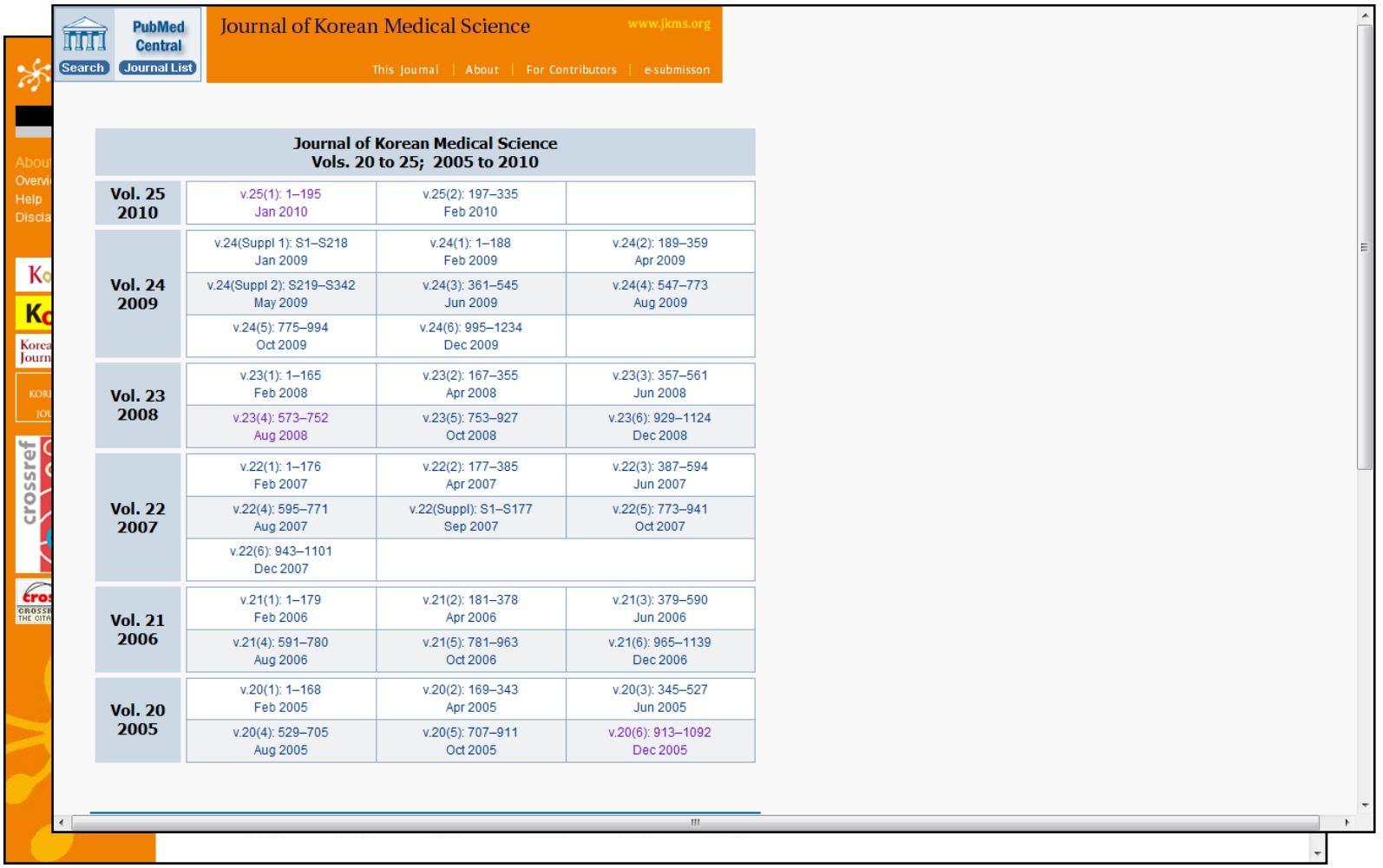

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2008년 SCI 논문들이 JKMS 논문을 인용한 횟수의 순위로 10위까지 열거함. Journal Citation Reports (JCR)의 가장 최근 출판일인 2009년 6월 30일에 조사한 기록을 담았으며, 1년마다 갱신함.

JKMS: Most Read articles

PMC에서 full-text와 PDF에 접속한 횟수에 기반하여 가장 많이 이용된 논문들에 대하여 순위를 매긴 것으로 분기별로 갱신한다. PMC에서 현재 JKMS는 2005년 부터만 검색되기 때문에 여기서의 순위는 PMC에 올라간 2005-2009년의 것에 한정되어 있다.

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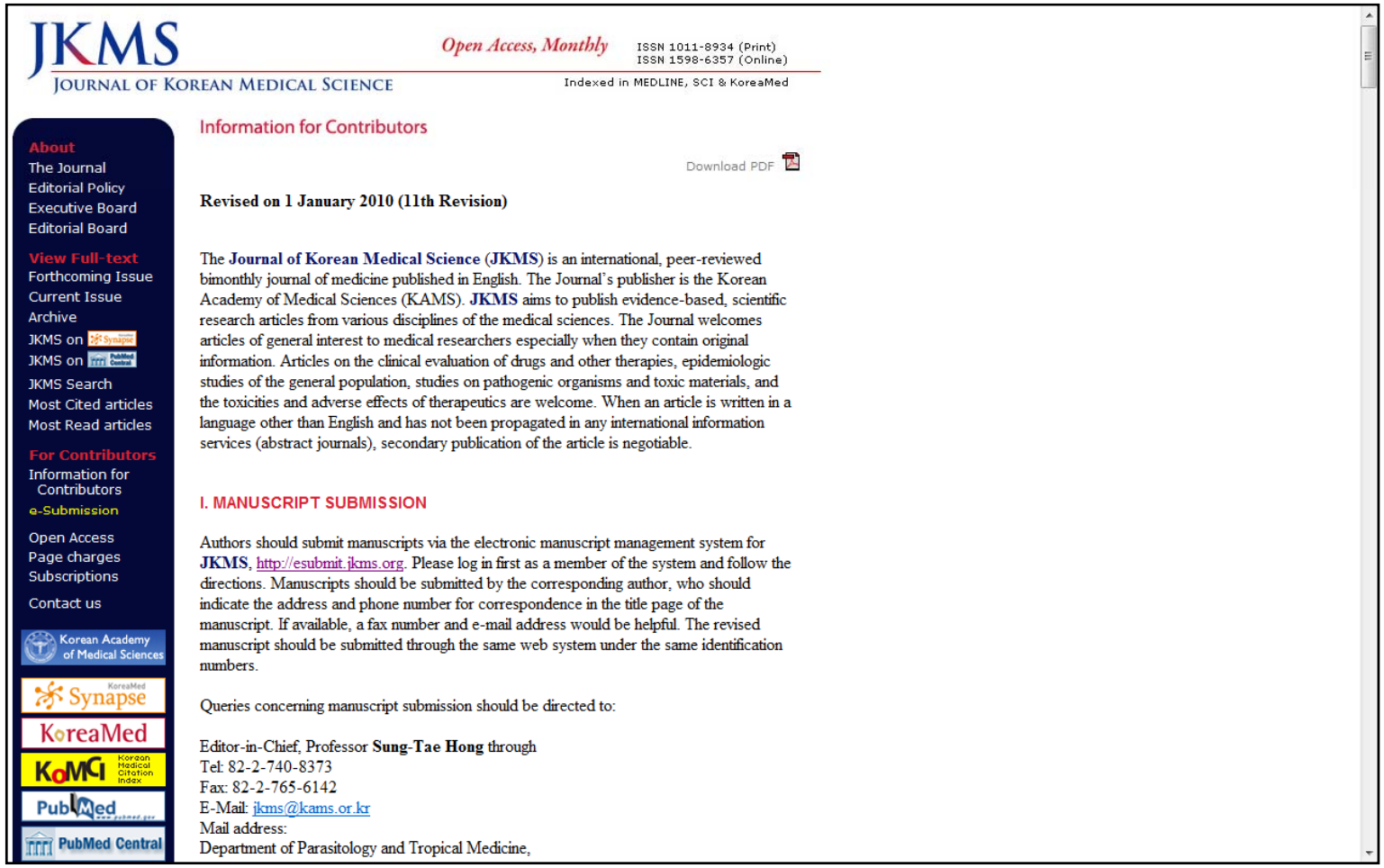

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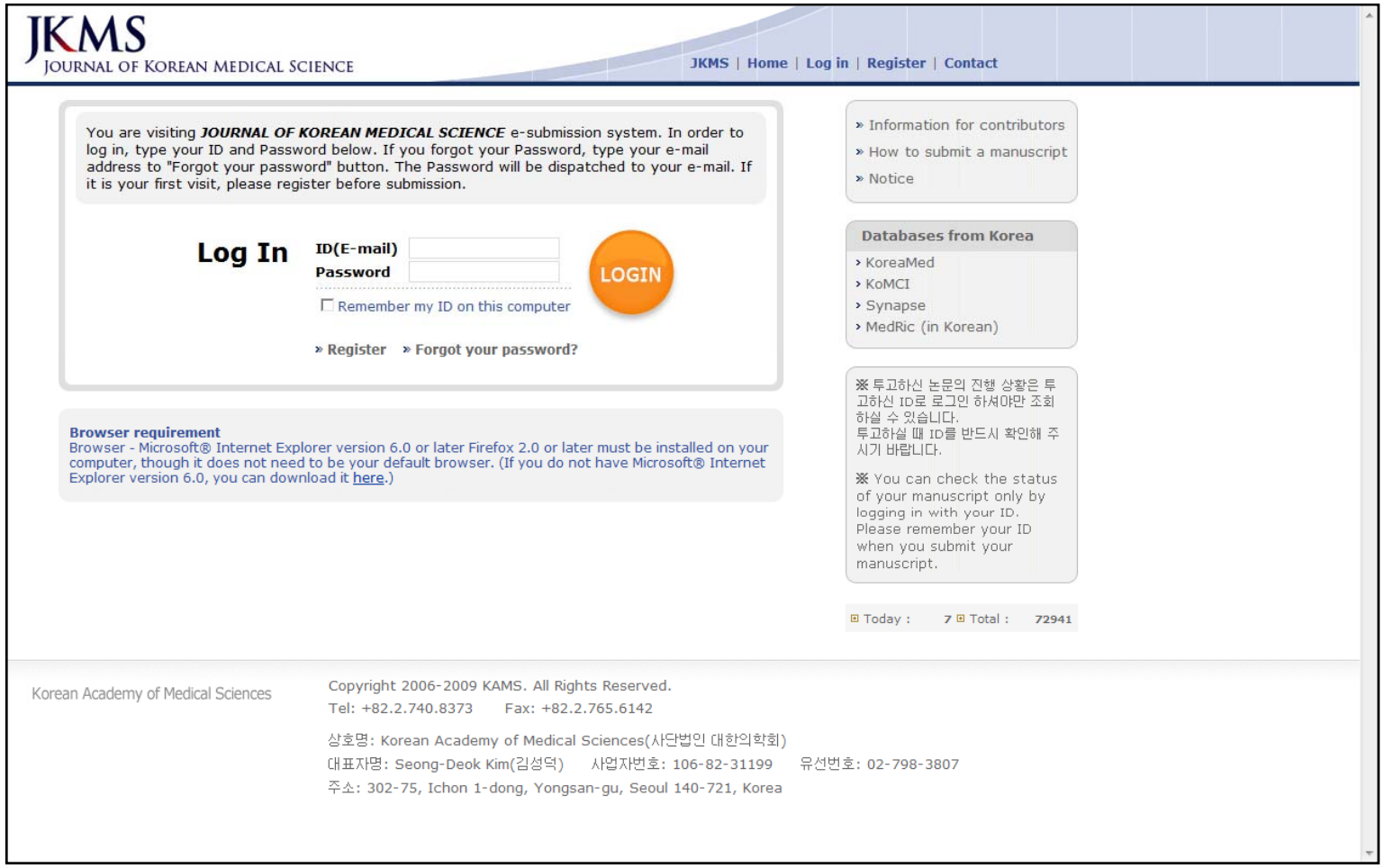

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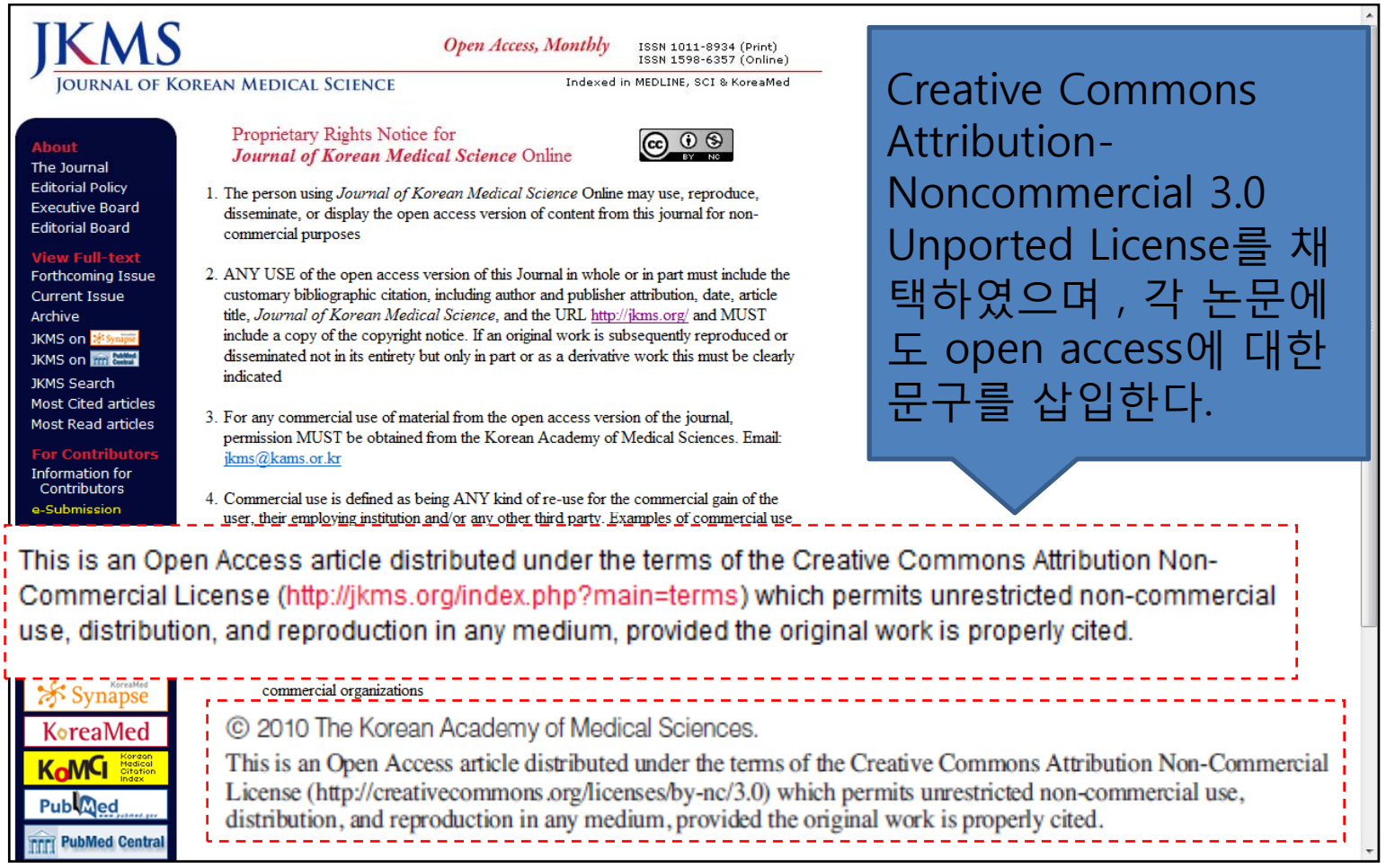

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