

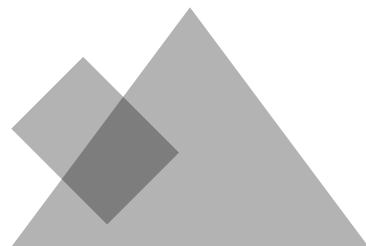
# PubMed: Using History

Prof Oluwadiya KS  
[www.oluwadiya.com](http://www.oluwadiya.com)

The logo for HINARI Research in Health is centered on the right side of the slide. It features the word "HINARI" in a large, blue, sans-serif font. Below it, the words "Research in Health" are written in a smaller, orange, sans-serif font. The logo is set against a light blue and white background that is part of a larger decorative graphic on the right side of the slide, which consists of several overlapping, semi-transparent, curved bands in shades of light blue and purple.



# Table of Contents

- History
  - Accessing full text articles from HINARI/PubMed
  - Full text article access problems
- 



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Search

- HINARI
- About HINARI
- Access the content
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- Partners
- Training materials
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Map of country breakdown  
 png, 122kb  
 More than 5000 institutions registered in 106 countries in 2012

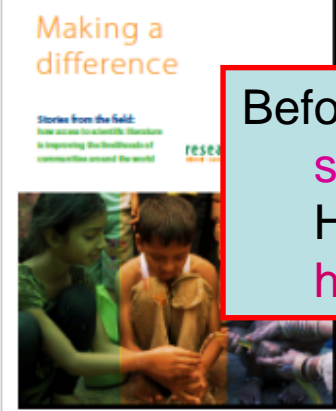
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 HINARI site in portuguese

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To celebrate Research4Life's 10th anniversary in 2011, we launched a user experience competition. We asked users to share with us how HINARI, AGORA or OARE has improved their work, life and community. In total...

Read more...

Before logging into the **Partner Publisher services** websites, we will **Login** to the HINARI site using the URL <http://www.who.int/hinari/>

OARE  
 RESEARCH IN THE ENVIRONMENT  
 Access to Research in the Environment  
 Brief video about OARE

News  
 HINARI and Research4Life programmes launch new portals

New Partners since the beginning of 2012  
 We are pleased to recognize the

ARDI

We will need to enter our HINARI **User Name** and **Password** in the appropriate boxes, then click on the **Login** button.

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Language:

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## PubMed

To access the **History** option, click on the **Advanced Search**

PubMed comprises more than 21 million citations for biomedical literature from MEDLINE, life science journals, and online books. Citations may include links to full-text content from PubMed Central and publisher web sites.

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### PubMed Advanced Search Builder

History deleted.

Use the builder below to create your search

[Edit](#)

#### Builder

	All Fields	<input type="text"/>	<input type="button" value="-"/>	<a href="#">Show index list</a>	
AND	All Fields	<input type="text"/>	<input type="button" value="-"/>	<input type="button" value="+"/>	<a href="#">Show index list</a>

or [Add to history](#)

#### History

[Clear history](#)

There is no recent history

We will discuss **History**, which is a feature of the **Advanced Search Builder** option.

Note that **Search History** will be lost after **8 hours** and the maximum number of searches is **100**.



PubMed  Search  
[Advanced](#) [Help](#)



# PubMed

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To build a search using **History**, begin by putting in your broadest search term. In this example, we will use *public health* as our main concept.



Choose additional filters

Text availability Abstract available Free full text available Full text available

Publication dates 5 years 10 years Custom range...

Species Humans Other Animals

Article types Clinical Trial Meta-Analysis Practice Guideline Randomized Controlled Trial Review Systematic Reviews more ...

Display Settings: Summary, 20 per page, Sorted by Recently Added Send to:

Filter your results: All (5071295) Free Full Text (847921) HINARI (1910867) Manage Filters

Results: 1 to 20 of 5071295 Prev Page 1 of 253565 Next > Last >>

- Misuse of prescription and illicit drugs among high-risk young adults in Los Angeles and New York... Association Between... Utilization of Primary Care Physician Services in Pennsylvania... (+)Equol inhibits invasion in prostate cancer DU145 cells possibly via down-regulation of matrix metalloproteinase-9, matrix metalloproteinase-2 and urokinase-type plasminogen activator by antioxidant activity...

The result for the public health search is over 5 million articles. This can be viewed as a search set by clicking on the Advanced Search (History) page.

Related searches bmc public health public health nursing public health nutrition public health surveillance public health policy

Titles with your search terms Public health: The toxic truth about sugar. [Nature. 2012]

### PubMed Advanced Search Builder

Use the builder below to create your search

[Edit](#)

#### Builder

All Fields

AND All Fields

[Search](#) or [Add to history](#)

On the **Advanced Search (History)** page, our first search is given a set number identified by the # symbol - in this example it is #1. On the right side of the page, the number of articles is shown under the **Items found** column. In this case, a total of over **5 million** citations. We will return to **PubMed Home** by clicking on the hypertext link.

Note that the numbers for your searches may vary depending if you have previously searched that day in PubMed.

#### History

[Clear history](#)

Search	Add to builder	Query	Items found	Time
<a href="#">#1</a>	<a href="#">Add</a>	Search public health	<a href="#">5071295</a>	5:04:49

Choose additional filters

Text availability Abstract available Free full text available Full text available

Publication dates 5 years 10 years Custom range...

Species Humans Other Animals

Article types Clinical Trial Meta-Analysis Practice Guideline Randomized Controlled Trial

Display Settings: Summary, 20 per page, Sorted by Recently Added

Send to:

Filter your results: All (62601) Free Full Text (17655) HINARI (22945) Manage Filters

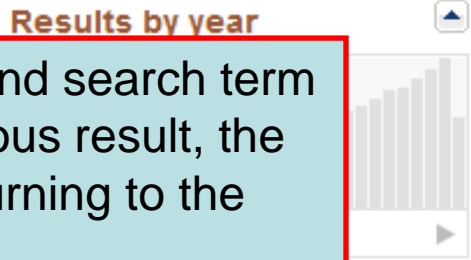
Results: 1 to 20 of 62601

1. Cochrane Column \* Rapid diagnostic tests can extend access of diagnostic services for uncomplicated Plasmodium falciparum malaria \* Commentary: Rapid diagnostic tests for diagnosing

We now have searched for malaria as a second search term and have 62601 citations. As in the previous result, the malaria search also can be viewed by returning to the Advanced Search (History) page.

Int J Epidemiol. 2012 Jun;41(3):607-10. No abstract available. PMID: 22798691 [PubMed - in process] Related citations

2. Iron and malaria interactions: research needs from basic science to global policy. Prentice AM, Cox SE. Adv Nutr. 2012 Jul 1;3(4):583-91. doi: 10.3945/an.111.001230. PMID: 22797996 [PubMed - in process] Related citations



Related searches cerebral malaria malaria vaccine severe malaria malaria pregnancy malaria review

# PubMed Advanced Search Builder

[YouTube Tutorial](#)

(#1) AND #2

[Edit](#)

[Clear](#)

## Builder

Recent Query #1

AND Recent Query #2

AND All Fields

[Search](#) or [Add to history](#)

We now will click on **#1 AND #2** to create a combined search for the two search terms – *public health AND malaria*. These two search terms also are added to the **Builder**. Note that there are two options – either **Search** or **Add to history** which will display the results on this page.

## History

[Clear history](#)

Search	Add to builder	Query	Items found	Time
#2	<a href="#">Add</a>	Search <b>malaria</b>	<a href="#">62601</a>	15:07:15
#1	<a href="#">Add</a>	Search <b>public health</b>	<a href="#">5071295</a>	15:04:49

Choose additional filters

Display Settings: Summary, 20 per page, Sorted by Recently Added

Filter your results:  
All (23631)  
Free Full Text (7979)  
HINARI (9842)  
Manage Filters

Text availability  
Abstract available  
Free full text available  
Full text available

Results: 1 to 20 of 23631

- 1. [Quantifying the indirect effects of key child survival interventions for pneumonia, diarrhoea, and measles.](#)  
Walker CL, Munos MK, Black RE.  
Epidemiol Infect. 2012 Jul 16:1-17. [Epub ahead of print]  
PMID: 22793874 [PubMed - as supplied by publisher]  
[Related citations](#)
- 2. [Using community health workers in community-based growth promotion: what stakeholders think.](#)  
Afulani P  
Health Ed  
PMID: 22793874 [PubMed - as supplied by publisher]  
[Related citations](#)
- 3. [Superintending the construction of a health facility: a case study from a rural area in South Africa.](#)  
Klein EY, Smith DL, Laxminarayan R, Levin S.  
Proc Biol Sci. 2012 Jul 11. [Epub ahead of print]  
PMID: 22787024 [PubMed - as supplied by publisher]  
[Related citations](#)



related data

base:

ct

Find items

Search details

(#1) AND #2

The search has narrowed the results down further to 23631 articles. Click on the Advanced link to return to the Advanced Search (History) page.

### PubMed Advanced Search Builder

(#3) AND asia

[Edit](#)

[Clear](#)

We now combine the latest search #3 with the geographic term Asia by adding both terms in the Builder fields. You can click on #3 and add Asia to the next line. We will use the Add to history option.

### Builder

Recent Query

AND

AND

[Show index list](#)

[Show index list](#)

[Search](#) or [Add to history](#)

We now combine the latest search #3 with the geographic term Asia.

### History

[Clear history](#)

Search	Add to builder	Query	Items found	Time
<a href="#">#3</a>	<a href="#">Add</a>	Search (#1) AND #2	<a href="#">23631</a>	15:13:14
<a href="#">#2</a>	<a href="#">Add</a>	Search malaria	<a href="#">62601</a>	15:07:15
<a href="#">#1</a>	<a href="#">Add</a>	Search public health	<a href="#">5071295</a>	15:04:49

### PubMed Advanced Search Builder

Use the builder below to create your search

[Edit](#)

[Clear](#)

#### Builder

All Fields

AND All Fields

[Search](#) or [Add to history](#)

Set #4 now is *public health AND malaria AND Asia* with a result of 4141 citations. Next we will open the searches' *Summary display setting* by clicking on the *Items Found #*.

Note: to clear these searches, click on *Clear History* button.

#### History

[Clear history](#)

Search	Add to builder	Query	Items found	Time
<a href="#">#4</a>	<a href="#">Add</a>	Search #3 AND asia	<a href="#">4141</a>	16:17:28
<a href="#">#3</a>	<a href="#">Add</a>	Search #1 AND #2	<a href="#">22657</a>	16:12:55
<a href="#">#2</a>	<a href="#">Add</a>	Search malaria	<a href="#">60453</a>	16:12:31
<a href="#">#1</a>	<a href="#">Add</a>	Search public health	<a href="#">4886747</a>	16:12:22

Choose additional filters

Text availability  
Abstract available  
Free full text available  
Full text available

Publication dates  
5 years  
10 years  
Custom range...

Species  
Humans  
Other Animals

Article types  
Clinical Trial  
Meta-Analysis  
Practice Guideline  
Randomized Controlled Trial  
Review  
Systematic Reviews  
more ...

Display Settings:  Summary, 20 per page, Sorted by Recently Added

Send to:

Filter your results:  
All (4309)  
Free Full Text (1161)  
HINARI (1275)  
Manage Filters

See 19 articles found by citation matching your search:  
[Distribution of organochlorine pesticides \(OCs\) and polychlorinated biphenyls \(PCBs\) in marine sediments directly exposed to wastewater from Cortiou, Marseille, Syak](#)  
[Aliphatic hydrocarbons, polycyclic aromatic hydrocarbons, biphenyls, organochlorine, and organophosphorus compounds in sediments from the Arc river estuary, France](#)  
[al. Environ Sci Pollut Res Int.](#)  
[Research in conflict zones: implications for the health of women](#)  
[al. ANS Adv Nurs Sci. \(2010\)](#)

This is the Summary display for the combined #4 search.  
We now will proceed to the accessing full-text articles slides.

Results: 1 to 20 of 4309 << First < Prev Page 1 of 216 Next > Last >>

- [Two functional reticulocyte binding-like \(RBL\) invasion ligands of zoonotic Plasmodium knowlesi exhibit differential adhesion to monkey and human erythrocytes.](#)  
Semenya AA, Tran TM, Meyer EV, Barnwell JW, Galinski MR. Malar J. 2012 Jul 6;11(1):228. [Epub ahead of print]  
PMID: 22770469 [PubMed - as supplied by publisher] **Free Article**  
[Related citations](#)
- [Analysis of Plasmodium falciparum diversity in natural infections by deep sequencing.](#)  
Manske M, Miotto O, Campino S, Auburn S, Almagro-Garcia J, Maslen G, O'Brien J, Djimde A, Doumbo O, Zongo I, Ouedraogo JB, Michon P, Mueller I, Siba P, Nzila A, Borrmann S, Kiara SM, Marsh K,

Find related data  
Database:  
Select  
Find items

Search details  
(#3) AND ("asia"[MeSH Terms] OR "asia"[All Fields])





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- [Topic-Specific Queries](#)

## More Resources

- [Clinical Trials](#)
- [E-Utilities](#)
- [LinkOut](#)

We now will discuss accessing full-text articles. Enter the search *chloroquine resistance*.

[Choose additional filters](#)

**Text availability**  
Abstract available  
Free full text available  
Full text available

**Publication dates**  
5 years  
10 years  
Custom range...

**Species**  
Humans  
Other Animals

**Article types**  
Clinical Trial  
Meta-Analysis  
Practice Guideline  
Randomized Controlled Trial  
Review  
Systematic Reviews  
more ...

**Display Settings:**  Summary, 20 per page, Sorted by Recently Added

**Send to:**

**Filter your results:**  
All (4263)  
[Free Full Text \(1170\)](#)  
[HINARI \(1537\)](#)  
[Manage Filters](#)

**Results: 1 to 20 of 4263** < First < Prev Page 1 of 214 Next > Last >>

- [Blockage of Spontaneous Intraerythrocytic Plasmodium](#)  
1. Enomoto M, Kawazu S, J, Mikoshiba K. PLoS One. 2012;7(7):e3949. PMID: 22792177 [PubMed - [Related citations](#)]
- [Optimizing treatments for lymphangiomyomatosis.](#)  
2. Taveira-Dasilva AM, Moss J. Expert Rev Respir Med. 2012 Jun;6(3):267-76. PMID: 22788941 [PubMed - in process] [Related citations](#)
- [Optimization of 4-Aminoquinoline/Clotrimazole-Based Hybrid Antimalarials: Further Structure-Activity Relationships, in vivo Studies, and Preliminary Toxicity Profiling.](#)  
3. Gemma S, Camodeca C, Sanna Coccone S, Joshi BP, Bernetti M, Moretti V, Brogi S, Bonache MC, Savini L, Taramelli D, Basilico N, Parapini S, Rottmann M, Brun R, Lamponi S, Caccia S, Guiso G, Summer R, Martin R, Saponara S, Gorelli B, Novellino E, Campiani G, Butini S. J Med Chem. 2012 Jul 12. [Epub ahead of print] PMID: 22783984 [PubMed - as supplied by publisher] [Related citations](#)

The results for this search are displayed in the **Summary** format with a total of **4263** citations.

**Related searches**  
[chloroquine resistance pfcr](#)  
[plasmodium falciparum chloroquine resistance transporter](#)  
[chloroquine resistance plasmodium vivax](#)  
[malaria chloroquine resistance](#)  
**PMC Images search for chloroquine resistance**

Choose additional filters

Text availability Abstract available Free full text available Full text available

Publication dates 5 years 10 years Custom range...

Species Humans Other Animals

Article types Clinical Trial Meta-Analysis Practice Guideline Randomized Controlled Trial Review Systematic

Display Settings: Summary, 20 per page, Sorted by Recently Added

Filter your results:

Format	Items per page	Sort by
<input type="radio"/> Summary	<input type="radio"/> 5	<input checked="" type="radio"/> Recently Added
<input type="radio"/> Summary (text)	<input type="radio"/> 10	<input type="radio"/> Pub Date
<input checked="" type="radio"/> Abstract	<input checked="" type="radio"/> 20	<input type="radio"/> First Author
<input type="radio"/> Abstract (text)	<input type="radio"/> 50	<input type="radio"/> Last Author
<input type="radio"/> MEDLINE	<input type="radio"/> 100	<input type="radio"/> Journal
<input type="radio"/> XML	<input type="radio"/> 200	<input type="radio"/> Title
<input type="radio"/> PMID List		

We will change the display to Abstract from the Display Settings drop down menu.

PMID: 22792177 [PubMed - in process] Related citations

- Optimizing treatments for lymphangiomyomatosis.
- Taveira-Dasilva AM, Moss J. Expert Rev Respir Med. 2012 Jun;6(3):267-76. PMID: 22788941 [PubMed - in process] Related citations
- Optimization of 4-Aminoquinoline/Clotrimazole-Based Hybrid Antimalarials: Further Structure-Activity Relationships, in vivo Studies, and Preliminary Toxicity Profiling. Gemma S, Camodeca C, Sanna Coccone S, Joshi BP, Bernetti M, Moretti V, Brogi S, Bonache MC, Savini L, Taramelli D, Basilico N, Parapini S, Rottmann M, Brun R, Lamponi S, Caccia S, Guiso G, Summer R, Martin R, Saponara S, Gorelli B, Novellino E, Campiani G, Butini S.

(1170)

Manage Filters

Related searches

- chloroquine resistance pfcr
- plasmodium falciparum chloroquine resistance transporter
- chloroquine resistance plasmodium vivax
- malaria chloroquine resistance

PMC Images search for

8. **Molecular assessment of Plasmodium falciparum resistance to antimalarial drugs in China.**

Zhang GQ, Guan YY, Zheng B, Wu S, Tang LH.

National Institute of Parasitic Diseases, Chinese Center for Disease Control and Prevention, WHO Collaborating Center for Malaria, Schistosomiasis and Filariasis, Shanghai 200025, China.

**OBJECTIVE:** In China, Chloroquine (CQ) and sulfadoxine-pyrimethamine (SP) were abandoned for the treatment of falciparum malaria 20 years ago due to resistance. Subsequent field studies showed a trend of declining CQ and SP resistance in the country. The main purpose of this study was to analyse the molecular markers of antimalarial resistance and thereby to assess the possibility of reintroduction of CQ or SP for falciparum malaria treatment. **METHODS:** Plasmodium falciparum field isolates were collected in 2006-2007 from Hainan and Yunnan provinces, China. Nested PCR-sequencing assays were applied to analyse the SNPs in four genes: P. falciparum chloroquine resistance transporter (pfcr1) gene, multi-drug resistance 1 (pfmdr1) gene, dihydrofolate reductase (dhfr) gene and dihydropteroate synthetase (dhps) gene. **RESULTS:** We found the widespread presence of point mutations in the dhfr and dhps genes which are associated with SP treatment failure. The molecular analyses also showed the fairly high prevalence of point mutation in the pfcr1 gene which is linked to CQ resistance. **CONCLUSION:** The results of the present study indicate that CQ and SP should not be reintroduced for falciparum malaria treatment in the near future in China.

PMID: 19772548 [PubMed - in process]

[Related articles](#)



[+ Publication Types](#)

From the **Abstract** display, links to full text publishers resources are shown at the bottom of the record.

9. **Hybrid drugs for malaria.**

Walsh JJ, Bell A.

Panoz Institute, Trinity College Dublin, Ireland. jjwalsh@tcd.ie

Malaria continues to devastate much of the tropics and sub-tropics in spite of the availability of a number of antimalarial drugs. Part of this problem is due to the disadvantages of the drugs in use, which include (depending on the drug) side effects, reduced efficacy due to resistance, and high cost. Multiple traditional and novel approaches to the discovery and design of new antimalarial agents are likely to be

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THE AMERICAN JOURNAL OF  
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Am. J. Trop. Med. Hyg., 80(2), 2009, pp. 215-217  
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**SHORT REPORT**

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If properly authenticated, you will see a URL that begins with:  
**hinari-gw.who.int/whalecommwww...**

**ABSTRACT**

The number of *Plasmodium vivax* malaria patients in the Republic of Korea and North Korea since the re-emergence of malaria in 1993 is estimated to be approximately one million. To cope with this situation, the Army of the Republic of Korea has performed chemoprophylaxis with hydroxychloroquine and primaquine since 1997. The cumulative number of soldiers in the Army of the Republic of Korea given chemoprophylaxis exceeded 1.4 million by 2007. Extensive chemoprophylaxis contributed to preventing a rapid increase of malaria patients in the Army of the Republic of Korea, but increased the possibility of the occurrence of chloroquine (CQ)-resistant *P. vivax* strains. In this study, treatment responses of *P. vivax* malaria patients in the Republic of Korea monitored during 2003–2007, and CQ resistance was confirmed in 2 of 484 enrolled patients. Our results are the first report of CQ-resistant *P. vivax* in a temperate region of Asia. Continuous surveillance is warranted to monitor the change in CQ resistance frequency of *P. vivax* in the Republic of Korea.

**QUICK SEARCH:** [advanced]

Author: Keyword(s):

Go

Year: Vol: Page:

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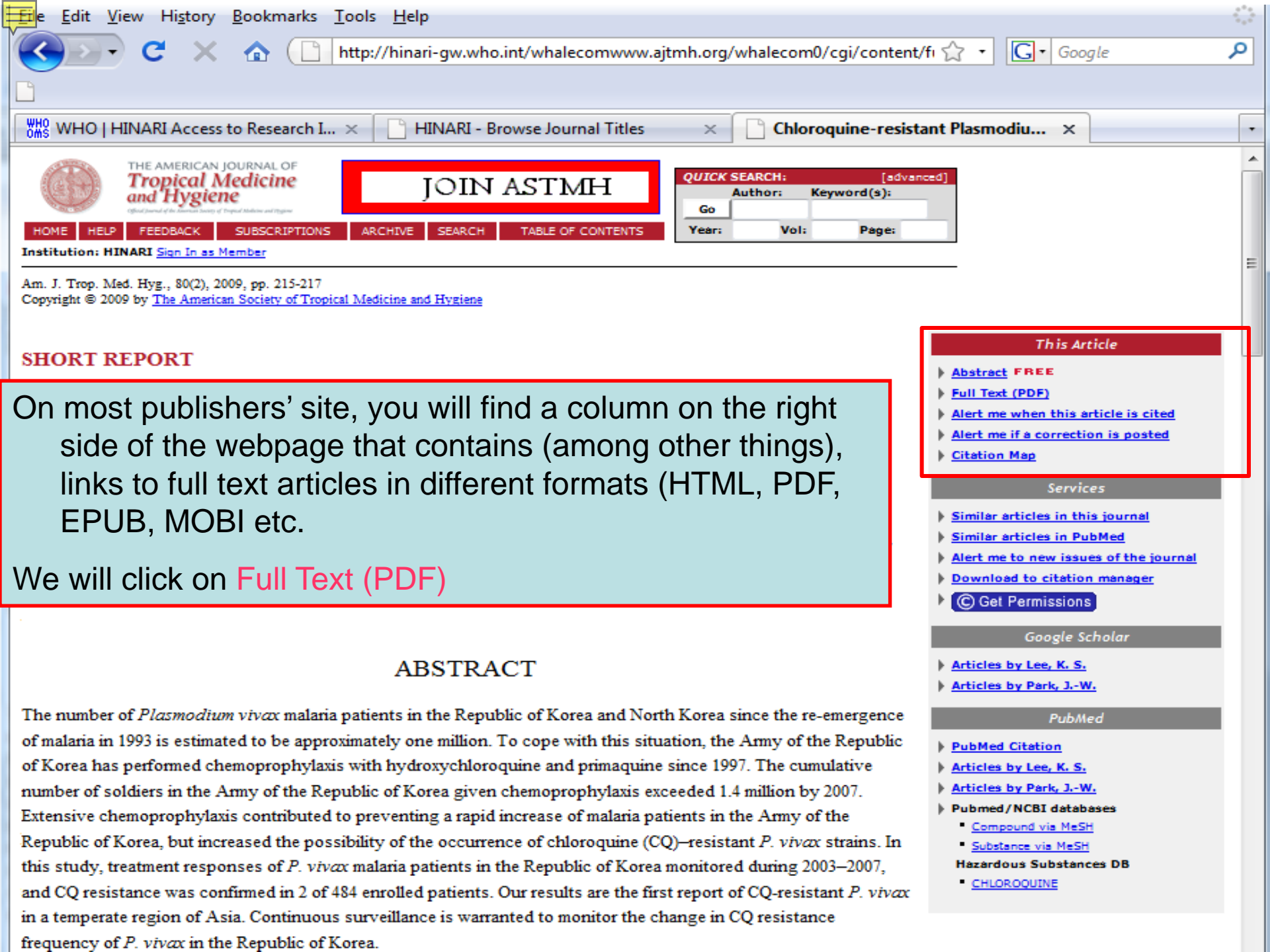
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- ▶ [Articles by Park, J.-W.](#)
- ▶ **PubMed/NCBI databases**
  - [Compound via MeSH](#)
  - [Substance via MeSH](#)
- ▶ **Hazardous Substances DB**
  - [CHLOROQUINE](#)



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Go	Author:	Keyword(s):
	<input type="text"/>	<input type="text"/>
Year:	Vol:	Page:
<input type="text"/>	<input type="text"/>	<input type="text"/>

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**SHORT REPORT**

On most publishers' site, you will find a column on the right side of the webpage that contains (among other things), links to full text articles in different formats (HTML, PDF, EPUB, MOBI etc).  
We will click on **Full Text (PDF)**

**ABSTRACT**

The number of *Plasmodium vivax* malaria patients in the Republic of Korea and North Korea since the re-emergence of malaria in 1993 is estimated to be approximately one million. To cope with this situation, the Army of the Republic of Korea has performed chemoprophylaxis with hydroxychloroquine and primaquine since 1997. The cumulative number of soldiers in the Army of the Republic of Korea given chemoprophylaxis exceeded 1.4 million by 2007. Extensive chemoprophylaxis contributed to preventing a rapid increase of malaria patients in the Army of the Republic of Korea, but increased the possibility of the occurrence of chloroquine (CQ)-resistant *P. vivax* strains. In this study, treatment responses of *P. vivax* malaria patients in the Republic of Korea monitored during 2003-2007, and CQ resistance was confirmed in 2 of 484 enrolled patients. Our results are the first report of CQ-resistant *P. vivax* in a temperate region of Asia. Continuous surveillance is warranted to monitor the change in CQ resistance frequency of *P. vivax* in the Republic of Korea.

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Am. J. Trop. Med. Hyg., 80(2), 2009, pp. 215-217  
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### Short Report: Chloroquine-resistant *Plasmodium vivax* in the Republic of Korea

Kkot Sil Lee,† Tae Hyong Kim,† Eu Suk Kim, Hyeong-Seok Lim, Joon-Sup Yeom, Gyo Jun, and Jae-Won Park\*

Department of Internal Medicine, Kwandong University College of Medicine, Seoul, Republic of Korea; Department of Internal Medicine, Soonchunhyang University College of Medicine, Seoul, Republic of Korea; Department of Internal Medicine, Dongguk University College of Medicine, Seoul, Republic of Korea; Department of Pharmacology, Ulsan University College of Medicine, Seoul, Republic of Korea; Department of Internal Medicine, Kangbuk Samsung Hospital, Sungkyunkwan University School of Medicine, Seoul, Republic of Korea; Departments of Microbiology and Biochemistry, Graduate School of Medicine, Gachon University of Medicine and Science, Incheon, Republic of Korea

**Abstract.** The number of *Plasmodium vivax* malaria patients in the Republic of Korea and North Korea since the re-emergence of malaria in 1993 is estimated to be approximately one million. To cope with this situation, the Army of the Republic of Korea has performed chemoprophylaxis with hydroxychloroquine and primaquine since 1997. The cumulative number of soldiers in the Army of the Republic of Korea given chemoprophylaxis exceeded 1.4 million by 2007. Extensive chemoprophylaxis contributed to preventing a rapid increase of malaria patients in the Army of the Republic of Korea, but increased the possibility of the occurrence of chloroquine (CQ)-resistant *P. vivax* strains. In this study, treatment responses of *P. vivax* malaria patients in the Republic of Korea monitored during 2003–2007, and CQ resistance of *P. vivax* in a temperate region of the Republic of Korea were investigated. The frequency of *P. vivax* in

*Plasmodium vivax* malaria in the Republic of Korea and North Korea since the re-emergence in 1993 in the Republic of Korea and North Korea since the re-emergence likely approaches one million.<sup>1-4</sup> To cope with the situation, the Army of the Republic of Korea has performed chemoprophylaxis with hydroxychloroquine (HCQ) and presumptive anti-relapse therapy with primaquine since 1997.<sup>5</sup> The cumulative number of the soldiers in the Army of the Republic of Korea given chemoprophylaxis exceeded 1.4 million by 2007. This extensive chemoprophylaxis campaign has helped prevent a rapid increase of malaria patients in the Army of the Republic of Korea. However, this success is tempered by the increased possibility of chloroquine (CQ)-resistant *P. vivax* strains.<sup>6</sup>

In this study, 484 patients from 6 hospitals in the Republic of Korea (5 in the malaria-prevalent region and 1 in Seoul) were enrolled during 2003–2007. Blood samples were collected from all patients before HCQ treatment and 24 hours after completion of treatment. Treatment responses were monitored by investigation of fever clearance time and par-

had not been in malaria-prevalent areas in other nations during the two years prior to their present hospitalization.

Patient A was a 26-year-old man (civilian) who had been discharged from the military in May 1998. Chemoprophylaxis was not performed during his military service. He was admitted to hospital I located in Goyang, a malaria-prevalent area in Kyonggi Province, on July 30, 2003. *Plasmodium vivax* malaria was confirmed and he was administered 2,000 mg of HCQ over a three-day period. More specifically, on day 0, he was given 800 mg of HCQ, with doses of 400 mg administered 6 hours and 24 hours later (day 1), and 48 hours later (day 2). Despite administration of the first cycle of HCQ treatment, fever did not subside until day 6 and *P. vivax* trophozoites were evident in a peripheral blood smear obtained on day 6. Parasite density on day 0 (before the treatment) and day 3 (24 h after completion of HCQ treatment) were 3,500/μL and 300/μL, respectively. Gene amplification by species-specific primers for small subunit ribosomal RNA<sup>8</sup> showed that *Plasmodia* in the patient's peripheral blood was *P. vivax*.



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Hydroxychloroquine has been reported to be as active as CQ against malaria parasites.<sup>15,16</sup> and 400 mg of HCQ is the molar equivalent of 309.6 mg of HCQ base and 295.0 mg of CQ base. Therefore, a CQ concentration of 10 ng/mL in plasma, which is the minimum effective concentration against CQ-susceptible *P. vivax*, is equivalent to an HCQ concentration of 10.5 ng/mL of plasma. In this study, treatment with 2,000 mg of HCQ over a three-day period was not effective in 2 (0.4%) of 484 patients. For these two patients, plasma concentrations of HCQ 24 hours after completion of HCQ treatments were much higher than the minimum effective concentration of CQ against *P. vivax*.<sup>17</sup> For the 482 patients with successful therapeutic outcomes, the mean and the standard deviation of plasma concentrations of HCQ 24 hours after completion of HCQ treatments were 220 ng/mL and 121 ng/mL, respectively, which were not distinct from the two patients in whom HCQ treatment failed. This indicates that HCQ was absorbed and metabolized normally in the two patients, precluding the possibility that the treatment failure was caused by personal factors. In the two patients, parasitemias were reduced markedly, but not cleared, by HCQ administration. Patient A was cured by additional administration of HCQ; this success may have been the result of the infecting *P. vivax* being exposed to an increased trough concentration of HCQ for an extended period because of the cumulative dosage.

The present observations are the first report of CQ-resistant *P. vivax* from a temperate region of Asia. Surveillance activity should be strengthened to monitor the change of CQ susceptibility of *P. vivax* in the Republic of Korea.

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† These authors contributed equally to this work. ↕

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selective advantage great enough to become the dominant *P.falciparum* type in Guinea-Bissau. This is most likely due to the efficacy of high-dose chloroquine as used in Guinea-Bissau, combined with a loss of fitness associated with pfcr76T.

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15. Trop Med Int Health. 2009 Oct;14(10):1251-7. Epub 2009 Aug 25.

### 15. Monitoring for multidrug-resistance: a cross-sectional analysis of pyrimethamine resistance in Uíge, Angola

Menegon M, Pearce RJ, Inojosa WO, Roper C, Severini C.

Department of Infectious, Parasitic and Immunology

OBJECTIVES: To assess the extent of multidrug resistance in Uíge, Angola, and to test whether the dhfr triple mutant allele was present in the population.

Seventy-one samples of blood from patients in Uíge Hospital in 2004 were screened for multidrug resistance.

RESULTS: Mutations in pfcr76 (codon 436, 437) were common. Amongst individuals with complete genetic information 13.7% carried all seven of these mutations. Flanking microsatellite analysis revealed the triple mutant pfdhfr was derived from the southeast Asian lineage, while the N51I+S108N double mutant pfdhfr alleles are a local origin. pATPase6 mutations were rare and S769N was not found.

CONCLUSION: The parasite population of Uíge Angola has high frequency mutations in pfcr76, dhfr and dhps associated with resistance to chloroquine and sulphadoxine pyrimethamine, reflecting past reliance on these two drugs which were the mainstay of treatment until recently. Our findings show that drug resistance in Uíge has occurred through a combination of local drug pressure and the regional and international dispersal of resistance mutant alleles.

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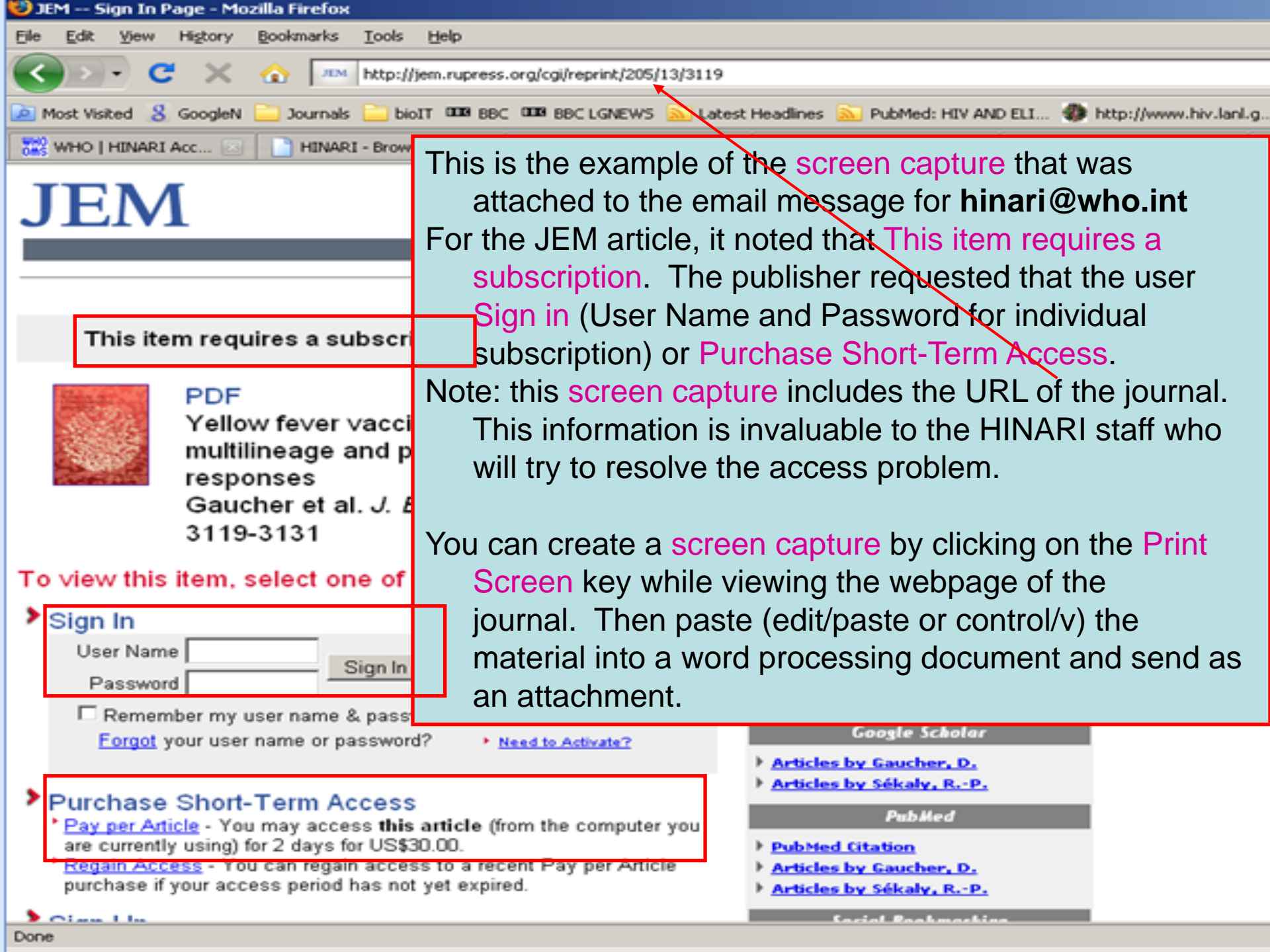
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