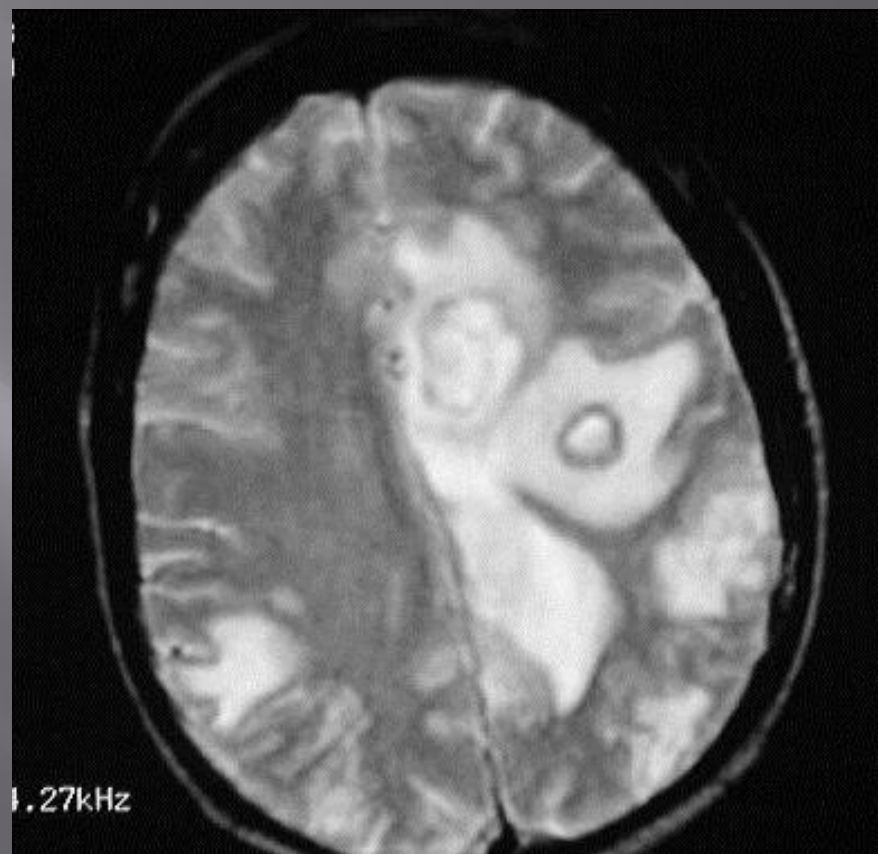
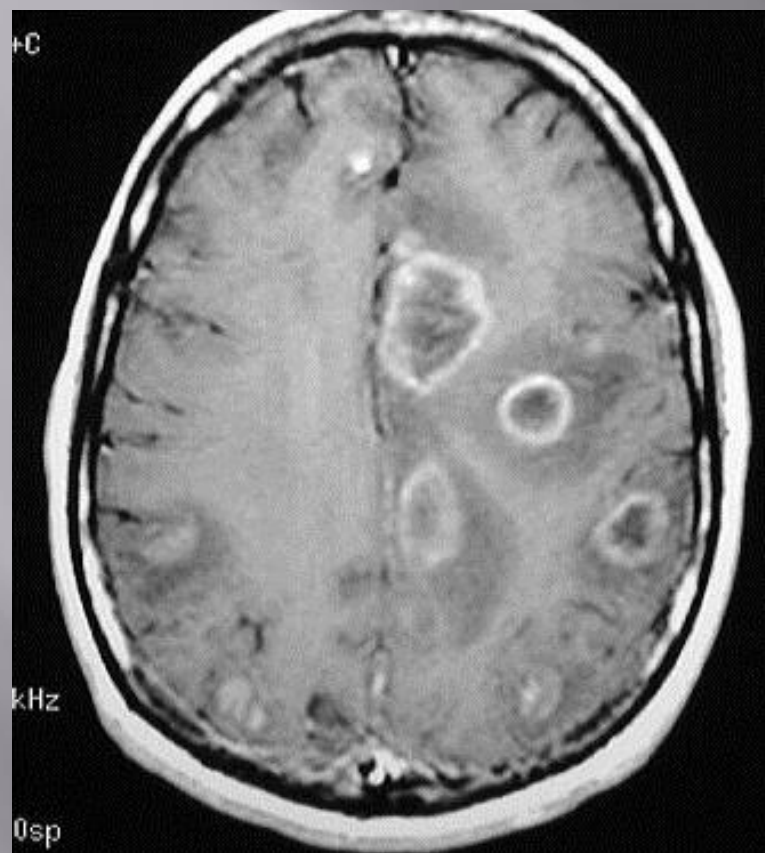


# PARAZİTER İNFEKSİYONLAR

## TOKSOPLAZMOZ

E. Ediz Tütüncü  
KLİMİK 2013

XVI. Türk Klinik Mikrobiyoloji ve  
İnfeksiyon Hastalıkları Kongresi  
15 Mart 2013, Antalya



Simple is beautiful.

## HASTA KONSÜLTASYON FORMU

Revizyon No: 01

Doküman No: YBH-FR-67

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Sayfa No: 1/1

Yayın Tarihi : 14.06.2006

İMVE  
sl

	<b>Konsültasyonu Öneren Şef/Uzmanın</b>	<b>Konsültasyonu İsteyen Doktorun</b>
	Adı Soyadı	Adı Soyadı
Erkek <input type="checkbox"/> Kadın <input type="checkbox"/>	İmza	İmza
	Birimi	Birimi
	Tarih/Saat	Tarih/Saat
	Konsültasyonun Yapılacağı Yer	
	Hasta Yatağında <input type="checkbox"/> Acil <input type="checkbox"/>	

Sa. Enf. Hast. Uzmanı,  
Toxo IgM (+) hastanın  
tarafından değerlendirilmiştir.





## Olgu

32 yaşında,  
12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok.

## Seroprevalence of Toxoplasmosis Among Pregnant Women in Kayseri

Tuba Kayman,<sup>1</sup> Mesut Kayman<sup>2</sup>

<sup>1</sup>Kayseri Eğitim ve Araştırma Hastanesi, Mikrobiyoloji Laboratuvarı, Kayseri

<sup>2</sup>Kayseri Doğumevi ve Çocuk Hastalıkları Hastanesi, Kadın Hastalıkları ve Doğum Kliniği, Kayseri

2006-2008  
Kayseri  
1676 gebe

*T. gondii* IgG (+) %33,9

## Erciyes Üniversitesi Tıp Fakültesi Hastanesi Parazitoloji Laboratuvarına Müracaat Eden Hastalarda *Anti-Toxoplasma gondii* Antikorlarının Dağılımı

Süleyman YAZAR \*  Salih KUK \* Ülfet ÇETİNKAYA \* Muhittin KAYA \* İzzet ŞAHİN \*

\* Erciyes Üniversitesi, Tıp Fakültesi, Tıbbi Parazitoloji Anabilim Dalı, TR-38039 Kayseri - TÜRKİYE

2009-2011

Kayseri

336 erkek, 1245 kadın,  
toplam 1581 kişi

*T. gondii* IgG (+) %28,8

**Gebelerde Toksoplasma gondii Seroprevalansı**  
*Seroprevalence of Toxoplasma gondii in Pregnant Women*

Servet Kölgeliler<sup>1</sup>, Hayati Demiraslan<sup>2</sup>, Bekir Katarş<sup>3</sup>, Dilek Güler<sup>3</sup>

<sup>1</sup>82. Yıl Devlet Hastanesi ve <sup>2</sup>Adıyaman Devlet Hastanesi Enfeksiyon Hastalıkları ve Klinik Mikrobiyoloji Kliniği, <sup>3</sup>Kadın Hastalıkları ve Doğum Kliniği-Adıyaman

2007-2008  
Adıyaman  
455 gebe

*T. gondii* IgG (+) %48,4





## Seroprevalence and Coinfections of *Toxoplasma gondii* in Childbearing Age Women in Turkey

\*I Akyar

Dept. of Microbiology, Medical Faculty, Acibadem University, Istanbul, Turkey

(Received 26 Jul 2010; accepted 25 Feb 2011)

### Abstract

**Background:** Our aim was to detect the rate of *Toxoplasma gondii* infections and the coinfections in childbearing age women in Turkey accompanying using seroprevalence data from a multicenter hospital setting.

**Methods:** Overall, 17751 childbearing age women through 16-45 years were included to the study between 2004 and 2010. The clinical samples of the patients were collected from 16 hospitals and medical centers mostly from Istanbul and three other cities from Turkey. Enzyme immunoassay tests were performed in our central laboratory in Istanbul to investigate *T. gondii* with other TORCH infections or Epstein Barr virus, Hepatitis B virus, Hepatitis C virus and Human Immunodeficiency virus as accompanying infections.

2004-2010

İstanbul, Bursa, Adana, Kayseri ve Kocaeli

16 merkez

17751 kadın

*T. gondii* IgG (+) %24,6



## **Seroprevalence and Coinfections of *Toxoplasma gondii* in Childbearing Age Women in Turkey**

*\*I Akyar*

*Dept. of Microbiology, Medical Faculty, Acibadem University, Istanbul, Turkey*

<b>Years</b>	<b><i>T. gondii</i> Ig M positive (n/%)</b>	<b><i>T. gondii</i> Ig G positive (n/%)</b>
<b>2004</b>	21/ 1.40	464/ 31.12
<b>2005</b>	21/ 1.15	798/ 43.77
<b>2006</b>	34/ 1.66	482/ 23.65
<b>2007</b>	36/ 1.22	602/ 20.52
<b>2008</b>	40/ 1.27	690/ 21.95
<b>2009</b>	55/ 1.41	829/ 21.38
<b>2010</b>	31/ 1.26	505/ 20.64
<b>Total</b>	238/ 1.34	4370/ 24.6

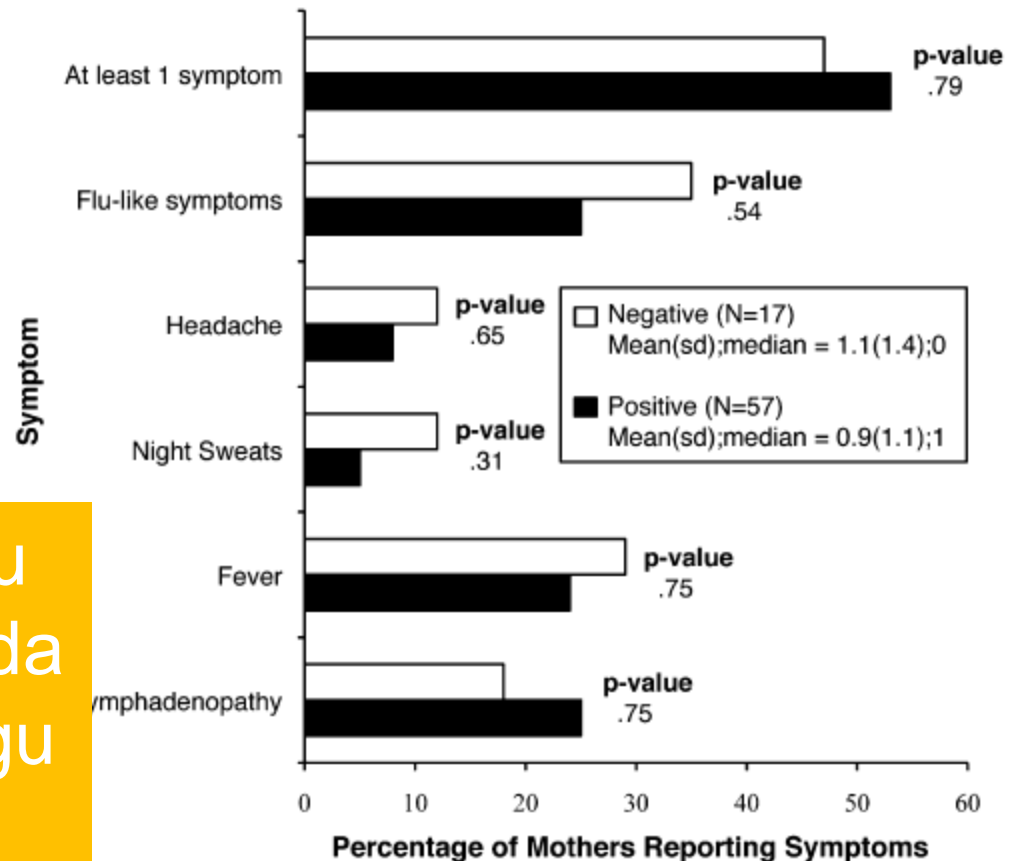


## Olgu

32 yaşında,  
12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok,  
Herhangi bir yakınması yok.

# Unrecognized Ingestion of *Toxoplasma gondii* Oocysts Leads to Congenital Toxoplasmosis and Causes Epidemics in North America

Kenneth Boyer,<sup>1,2</sup> Dolores Hill,<sup>3</sup> Ernest Mui,<sup>4</sup> Kristen Wroblewski,<sup>5</sup> Theodore Karrison,<sup>5</sup> J. P. Dubey,<sup>3</sup> Mari Sautter,<sup>4</sup> A. Gwendolyn Noble,<sup>4,6</sup> Shawn Withers,<sup>5</sup> Charles Swisher,<sup>7</sup> Peter Heydemann,<sup>1,8</sup> Tiffany Hosten,<sup>4</sup> Jane Babiarz,<sup>4</sup> Daniel Lee,<sup>4</sup> Paul Meier,<sup>5,9,\*</sup> Rima McLeod,<sup>4,10,11,12</sup> and other members of the Toxoplasmosis Study Group<sup>a</sup>



Akut *T. gondii* infeksiyonu geçiren gebelerin çoğunda belirgin semptom ve bulgu saptanmaz.

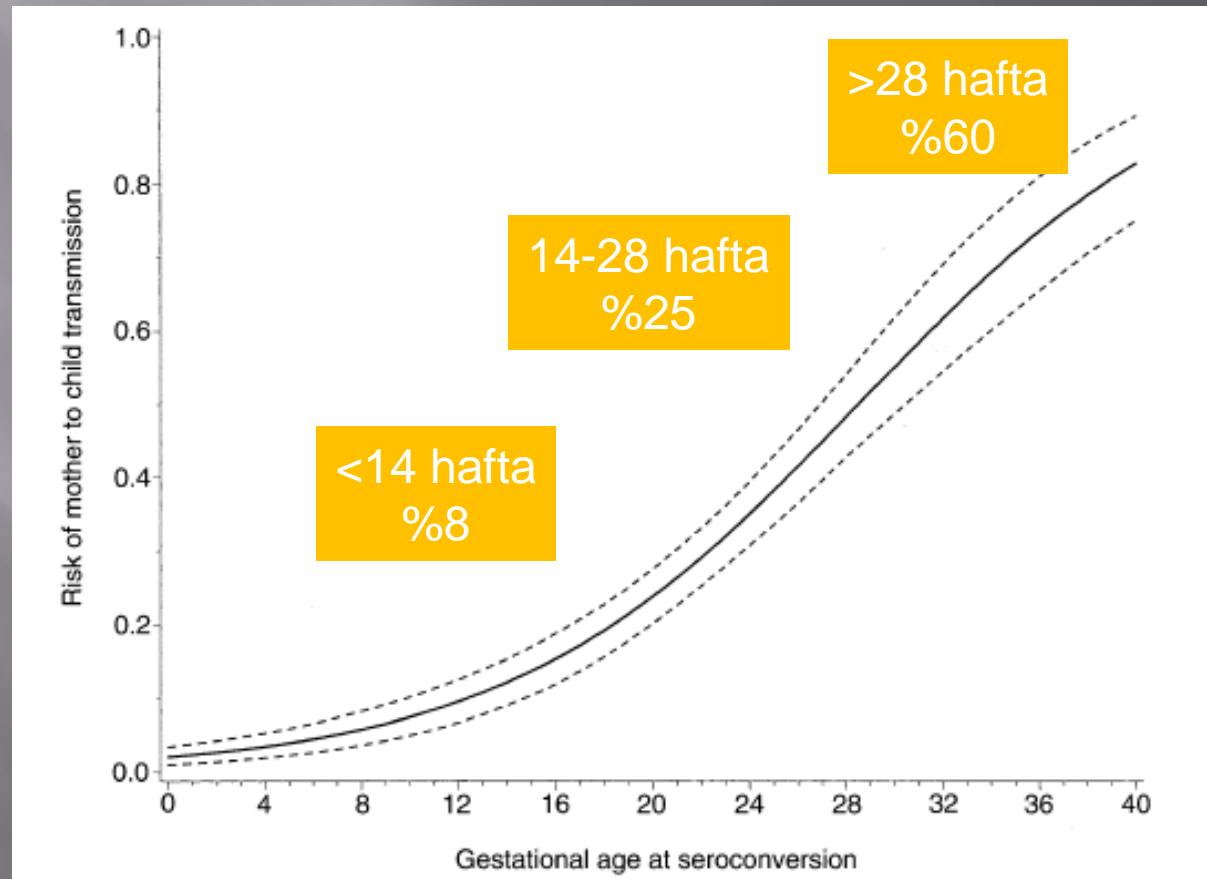
Konjenital toksoplazmoz tanısı alan çocuk sahibi olan annelerin %52'si, gebelikleri sırasında infeksiyon düşündürecek hiçbir tablo tanımlamamaktadırlar.

Primer maternal infeksiyon sırasında parazitlerin plasenta yoluyla geçişi fetal infeksiyona yol açar.  
maternal parazitemi,  
iyi gelişmiş plasental kan akımı,  
maternal bağışıklığın henüz gelişmemesi.

Parazitin plasentada inflamatuvar bir odak oluşturması ve gebeliğin ilerleyen dönemlerinde takizoitlerin fetal dolaşıma salınması.

## Effect of timing and type of treatment on the risk of mother to child transmission of *Toxoplasma gondii*

European Multicentre Study on Congenital Toxoplasmosis\*



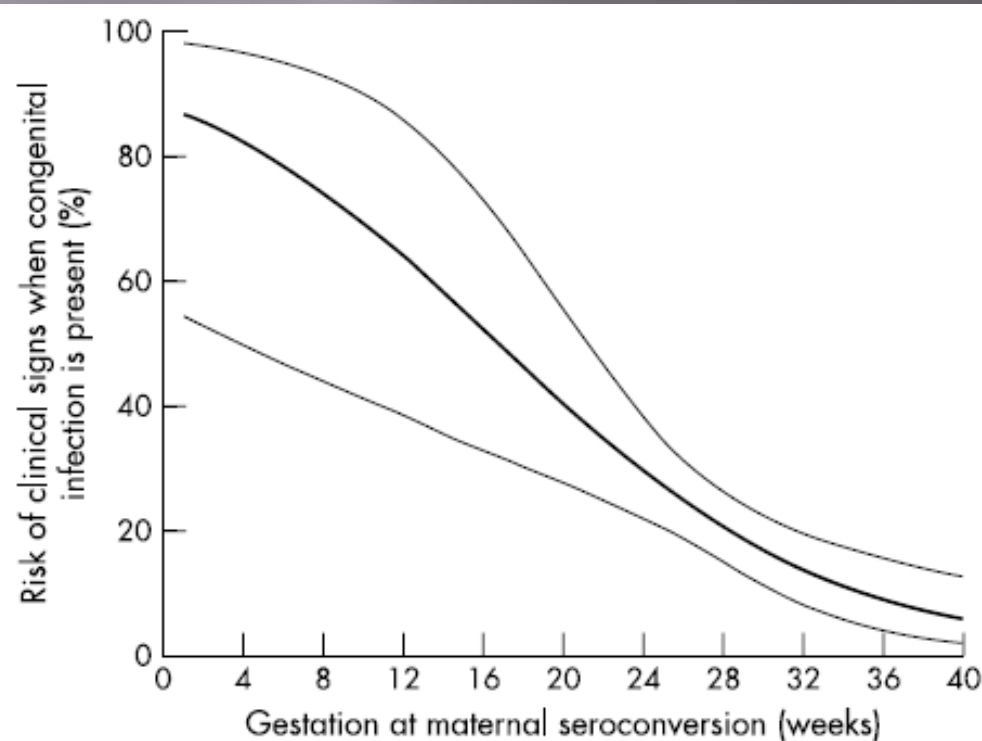
## REVIEW

# Congenital toxoplasmosis in the United Kingdom: to screen or not to screen?

R E Gilbert, C S Peckham

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*J Med Screen* 2002;**9**:135-141





## Management of *Toxoplasma gondii* Infection during Pregnancy

**Jose G. Montoya and Jack S. Remington**

Palo Alto Medical Foundation Toxoplasma Serology Laboratory, Palo Alto, and Department of Medicine and Division of Infectious Diseases and Geographic Medicine, Stanford University School of Medicine, Stanford, California

Tüm gebelerde, ideal olarak ilk trimesterde *T. gondii* IgG ve IgM ile serolojik tarama yapılması ve seronegatif kadınların izleme alınması optimal yaklaşımdır.



## Olgu

32 yaşında,  
12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok,  
Herhangi bir yakınması yok,  
*T. gondii* IgM (+).

Gebelik döneminde saptanan *T. gondii* IgM pozitifliği, her zaman yeni edinilmiş infeksiyon anlamına gelir mi?

## Duration of the IgM response in women acquiring *Toxoplasma gondii* during pregnancy: implications for clinical practice and cross-sectional incidence studies

L. GRAS<sup>1</sup>, R. E. GILBERT<sup>1\*</sup>, M. WALLON<sup>2</sup>, F. PEYRON<sup>2</sup>  
AND M. CORTINA-BORJA<sup>1</sup>

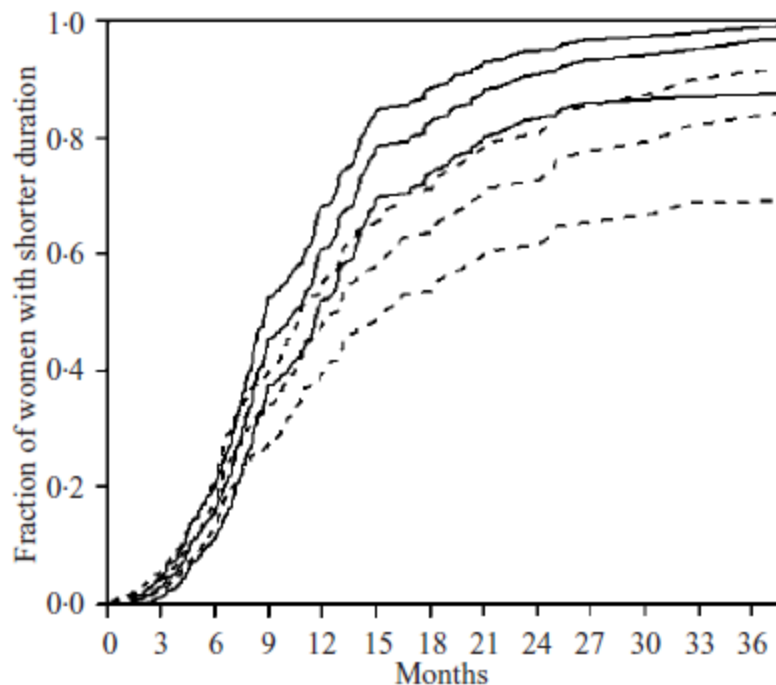


Fig. 1. Duration of detection of IgM antibodies and 95% CIs using the IFT (—), and the ISAGA test (---).

446 infekte gebede

ISAGA 12,8 ay median  
IFT 10,4 ay median

24. ayda pozitiflik  
ISAGA %27,1  
IFT %9,1

## Evaluation of Six Commercial Kits for Detection of Human Immunoglobulin M Antibodies to *Toxoplasma gondii*

MARIANNA WILSON,<sup>1\*</sup> JACK S. REMINGTON,<sup>2</sup> CHARLES CLAVET,<sup>3</sup> GEORGE VARNEY,<sup>3</sup> CYNTHIA PRESS,<sup>2</sup> DORIS WARE,<sup>1</sup> AND THE FDA TOXOPLASMOSIS AD HOC WORKING GROUP<sup>4†</sup>

*Division of Parasitic Diseases, National Center for Infectious Diseases, Centers for Disease Control and Prevention, Atlanta, Georgia<sup>1</sup>; Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto, California<sup>2</sup>; Winchester Engineering and Analytical Center, Food and Drug Administration, Winchester, Massachusetts<sup>3</sup>; and Center for Devices and Radiological Health, Food and Drug Administration, Rockville, Maryland<sup>4</sup>*

TABLE 2. Sensitivity, specificity, and rates of equivocal results for commercially available kits for detection of *Toxoplasma* IgM<sup>a</sup>

Kit	Sensitivity (%)	Specificity (%)	% Equivocal results
PAMF IgM ELISA	100	100 <sup>b</sup>	0
CDC EIA IgM	100	99.1	0
bioMérieux VIDAS Toxo IgM	100	98.6	0.9
Sanofi Platelia Toxo IgM	100	96.8	3.6
BioWhittaker Toxocap-M	100	95.9	2.7
Gull Toxo IgM	97	85.6	2.3
Abbott Toxo-M EIA	100	84.2	7.7
Abbott IMx Toxo IgM			
Version 1	100	77.5	9.0
Version 2	93.3	97.3	2.3

<sup>a</sup> Sensitivity was calculated on the basis of results for 30 samples, and specificity was calculated on the basis of results for 222 samples.

## Management of *Toxoplasma gondii* Infection during Pregnancy

**Jose G. Montoya and Jack S. Remington**

Palo Alto Medical Foundation Toxoplasma Serology Laboratory, Palo Alto, and Department of Medicine and Division of Infectious Diseases and Geographic Medicine, Stanford University School of Medicine, Stanford, California

IgM antikorları 12 aydan daha uzun süreyle saptanabileceği için, bir gebede pozitif ya da sınırda değer varlığı, referans laboratuvarlarda doğrulama gerektirir.

Pozitif IgM sonuçlarının ancak %40'ının gerçekten akut enfeksiyonu gösterdiği bildirilmiş.

## Management of *Toxoplasma gondii* Infection during Pregnancy

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IgM(+) / IgG(+)

Serum referans laboratuvara gönderilir.

IgM(+) / IgG(-)

Tetkik 1-3 hafta içinde tekrarlanmalıdır.

IgM(+) / IgG(-)

Klinik anlamı yok

IgM(+) / IgG(+)

Serokonversiyon



## Olgu

32 yaşında,  
12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok,  
Herhangi bir yakınması yok,  
*T. gondii* IgM (+),  
*T. gondii* IgG (+).





## Toksoplazma serolojik profili

Sabin-Feldman boya testi,  
AC/HS test,  
IgG avidite testi,  
IgA,  
IgE.

# Dyes as Microchemical Indicators of a New Immunity Phenomenon Affecting a Protozoon Parasite (*Toxoplasma*)<sup>1</sup>

ALBERT B. SABIN and HARRY A. FELDMAN<sup>2</sup>

*The Children's Hospital Research Foundation  
and Department of Pediatrics,  
University of Cincinnati College of Medicine*

The purpose of this preliminary communication is to describe a new immunity phenomenon in which dyes of certain chemical composition have been found capable of indicating the presence or absence of antibody activity. This phenomenon was discovered during the course of a search for some *in vitro* manifestation of the action of neutralizing antibody on toxoplasma, an obligate, intracellular protozoon parasite. After finding that toxoplasma in properly diluted mouse peritoneal exudate could be counted with great accuracy in a standard hemocytometer, we observed that in mixtures with immune serum the toxoplasma remained intact but lost the refractility they exhibited in mixtures with normal serum. When, after incubation at room temperature for several hours, large drops of such mixtures were allowed to dry slowly on slides overnight and then were stained with Wright's stain, large numbers of toxoplasma could be seen in the preparation from the normal serum mixture, whereas very few were found in that from the immune serum mixture. Small drops, spread thin and rapidly dried, revealed that, with few exceptions, the cytoplasm of the toxoplasma in the immune serum mixtures was distorted, poorly stained, or unstained as compared with the deep blue staining and granular structure of the cytoplasm of the toxoplasma in the mixtures with normal serum; the chromatin of the toxoplasma appeared the same in both types of mixtures. One of us (A. B. S.) had observed a number of years ago that, when alkaline methylene blue was added on a slide to a drop of peritoneal exudate containing toxoplasma, immediate deep purple staining of the parasites could be observed under the microscope. When this was done with

<sup>1</sup> Aided by a grant from the National Foundation for Infantile Paralysis.

<sup>2</sup> Senior Fellow, National Research Council.

Sabin- Feldman boya testi  
1948

# IS SABIN-FELDMAN DYE TEST USING *T. GONDII* TACHYZOITES FROM ANIMAL INOCULATION STILL THE BEST METHOD FOR DETECTING *TOXOPLASMA GONDII* ANTIBODIES?

Ruenruetai Udonsom<sup>1</sup>, Ruangrat Buddhirongawatr<sup>1,2</sup> and Yaowalark Sukthana<sup>1,3</sup>

<sup>1</sup>Department of Protozoology, Faculty of Tropical Medicine, Mahidol University, Bangkok; <sup>2</sup>Faculty of Veterinary Sciences, <sup>3</sup>International College, Mahidol University, Salaya Campus, Nakhon Pathom, Thailand

## Sabin- Feldman boya testi

Nötralizasyon testi,  
Canlı mikroorganizmalar kullanılır,  
IgG antikorlar saptanır,  
Duyarlı ve özgül,  
Altın standart

## Differential Agglutination Test for Diagnosis of Recently Acquired Infection with *Toxoplasma gondii*

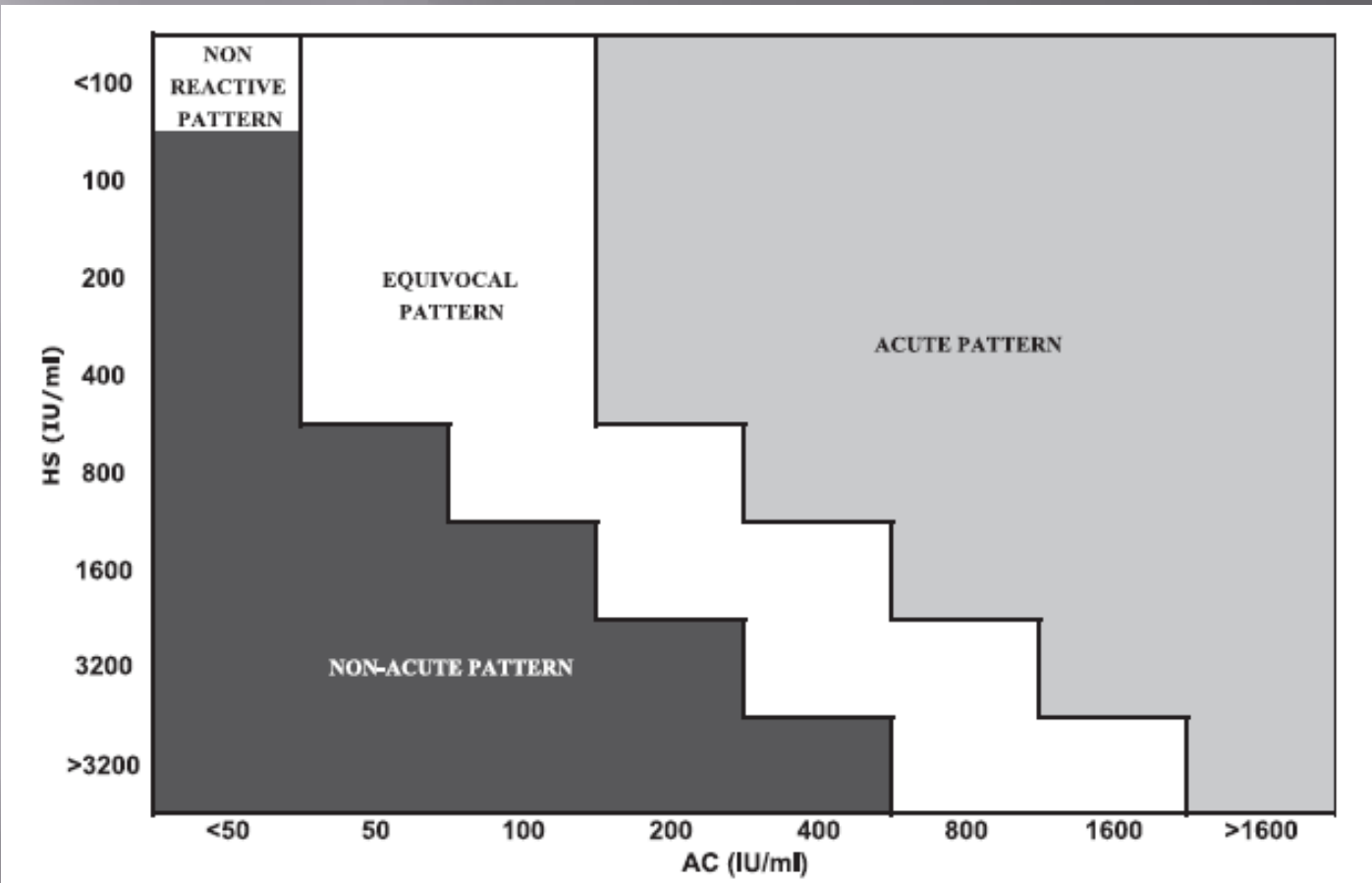
BRIAN R. DANNEMANN,<sup>1,2</sup> WINSTON C. VAUGHAN,<sup>2</sup> PHILLIPE THULLIEZ,<sup>3</sup> AND JACK S. REMINGTON<sup>1,2\*</sup>

*Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, 860 Bryant Street, Palo Alto, California 94301<sup>1</sup>; Stanford University School of Medicine, Stanford, California 94305<sup>2</sup>; and Laboratoire de Serologie Neonatale et de Recherche sur la Toxoplasmose, Institut de Puericulture de Paris, 75014 Paris, France<sup>3</sup>*

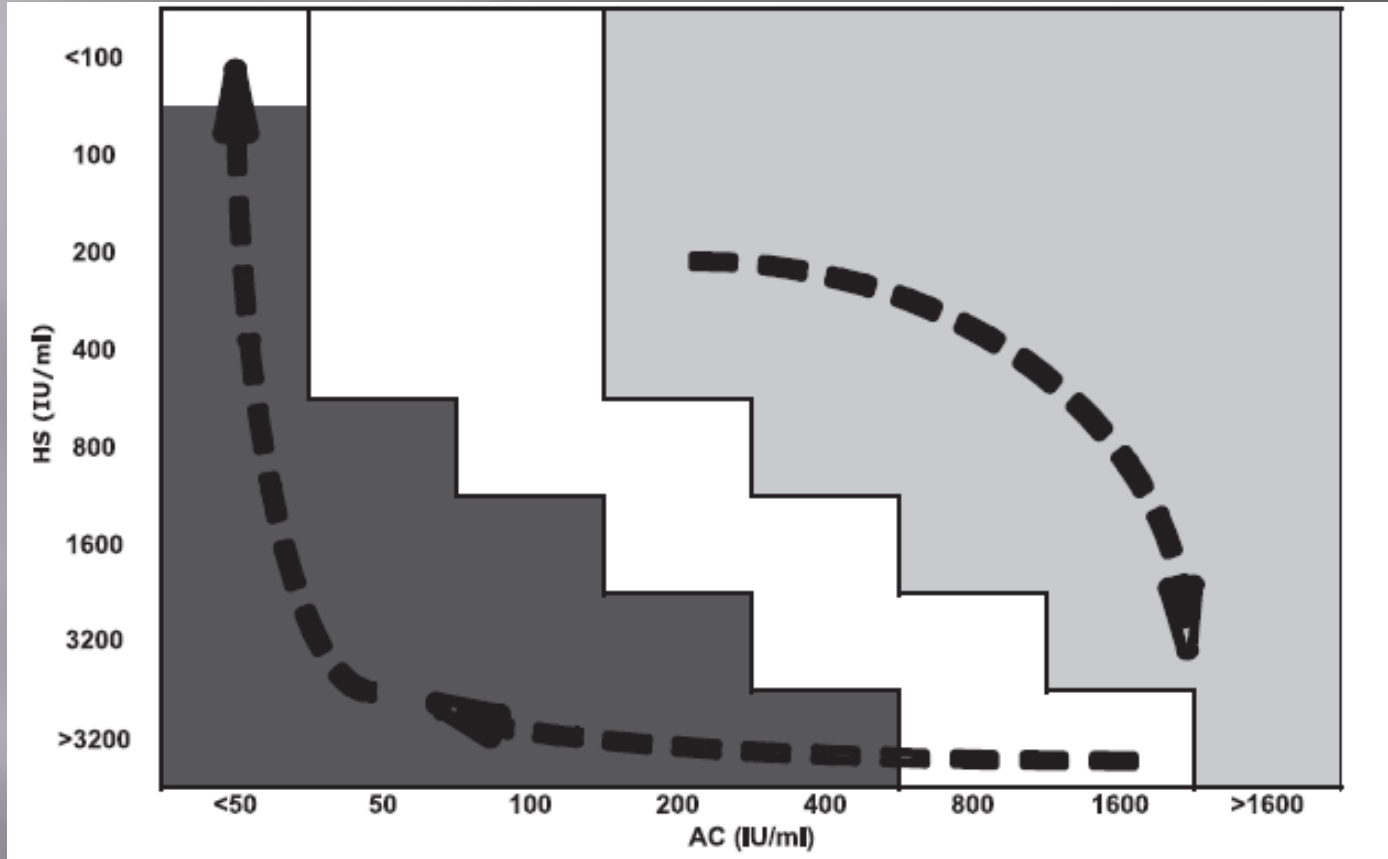
### AC/HS testi

Ayırt edici aglütinasyon testi,  
Aseton ve formalin ile muamele edilmiş  
takizoidlerin farklı aglütinasyon vermesi,

Akut dönemden kronik döneme ilerledikçe *T. gondii* yüzey antijenlerindeki değişikliklere bağlı olarak, farklı IgG profillerinin gelişmesi



Akut infeksiyonu olan hasta serumları hem AC hem de HS ile aglütinasyon verir, İnfeksiyonun üzerinden zaman geçtikçe AC titreleri düşer, HS yükselir.



Akut infeksiyonu olan hasta serumları hem AC hem de HS ile aglütinasyon verir, İnfeksiyonun üzerinden zaman geçtikçe AC titreleri düşer, HS yükselir.

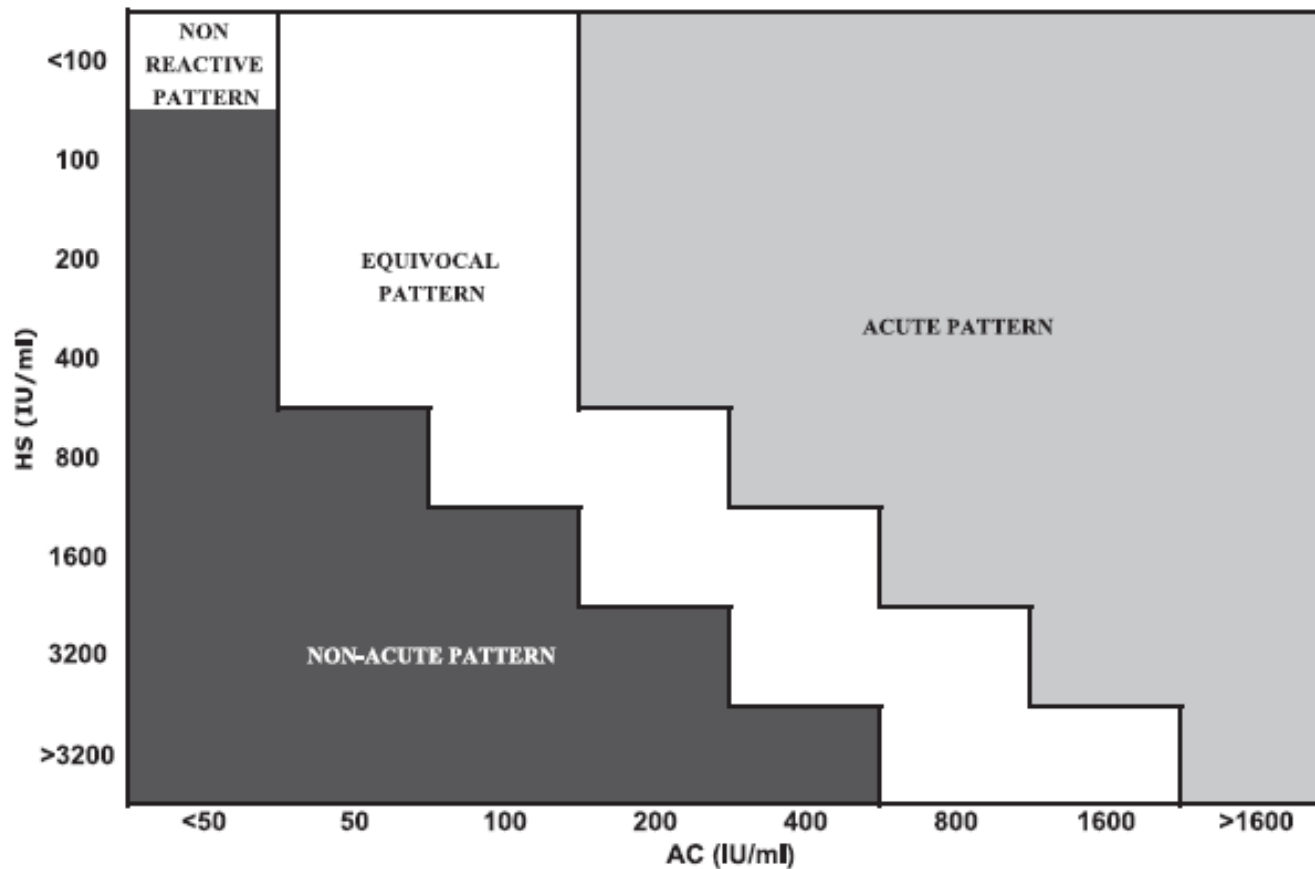


FIG. 1. Interpretative criteria for the differential agglutination (AC/HS) test.

AC/HS testi

Akut

Sınırdaki

Non-akut

Non-reaktif paternler



## The Differential Agglutination Test as a Diagnostic Aid in Cases of Toxoplasmic Lymphadenitis<sup>▽</sup>

Jose G. Montoya,<sup>1,2\*</sup> Andrew Berry,<sup>1</sup> Fernando Rosso,<sup>1,2</sup> and Jack S. Remington<sup>1,2</sup>

*Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto, California,<sup>1</sup> and  
Department of Medicine, Division of Infectious Diseases and Geographic Medicine,  
Stanford University School of Medicine, Stanford, California<sup>2</sup>*

Non-akut patern, akut infeksiyonun üzerinden en az 13 ay geçtiğini düşündürür.

Yüksek avidite testi ise akut infeksiyonun üzerinden 3-5 ay geçtiğini düşündürür.

13 ay sınırının gebelerde kullanımı konusunda çalışmalara ihtiyaç var...

## IgG avidite testi

Avidite=Fonksiyonel afinite,  
“Farklı” antikor popülasyonlarının  
antijeni bağlama gücünü ifade eder.

Spesifik IgG antikorlarının fonksiyonel  
afinitesi, başlangıçta düşüktür,  
B lenfosit seleksiyonu ile, izleyen  
haftalar-aylar içerisinde yükselir.

*T. gondii* ile akut infeksiyon sonrasında  
yüksek aviditeli antikorlar 3-5 ay içinde ortaya çıkar

## IgG avidite testi

üre gibi protein denatüre edici ajanlar kullanılarak, antijen-antikor kompleksleri ayrıştırılır.

Sonuçta, düşük aviditeli antikorlar tamamen çözünür/Ag'den ayrılırlarken, yüksek aviditeli antikorlar bağlı kalmaya devam ederler.

## Effect of Testing for IgG Avidity in the Diagnosis of *Toxoplasma gondii* Infection in Pregnant Women: Experience in a US Reference Laboratory

Oliver Liesenfeld,<sup>1,2,3</sup> Jose G. Montoya,<sup>1,2</sup>  
Sandra Kinney,<sup>1</sup> Cynthia Press,<sup>1</sup>  
and Jack S. Remington<sup>1,2</sup>

<sup>1</sup>Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto, and <sup>2</sup>Division of Infectious Diseases and Geographic Medicine, Department of Medicine, Stanford University School of Medicine, Stanford, California; <sup>3</sup>Department of Medical Microbiology and Immunology of Infection, Institute for Infection Medicine, Benjamin Franklin Medical Center, Free University of Berlin, Berlin, Germany

**Table 3.** Comparison of IgG avidity test and differential agglutination (AC/HS) test results in 125 serum samples taken from pregnant women during the first trimester.

Avidity result	AC/HS test result		
	Acute (n = 33)	Equivocal (n = 53)	Nonacute (n = 39)
Low	17 (51.5)	4 (7.5)	0
Borderline	12 (36.4)	18 (34.0)	2 (5.1)
High	4 (12.1)	31 (58.5)	37 (94.9)

NOTE. Data are no. (%) of serum samples. For further explanation of the AC/HS test, see figure 1.

AC/HS testi ile IgG avidite testi sonuçları birbirleriyle örtüşmektedir.

## Effect of Testing for IgG Avidity in the Diagnosis of *Toxoplasma gondii* Infection in Pregnant Women: Experience in a US Reference Laboratory

Oliver Liesenfeld,<sup>1,2,3</sup> Jose G. Montoya,<sup>1,2</sup>  
Sandra Kinney,<sup>1</sup> Cynthia Press,<sup>1</sup>  
and Jack S. Remington<sup>1,2</sup>

<sup>1</sup>Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto, and <sup>2</sup>Division of Infectious Diseases and Geographic Medicine, Department of Medicine, Stanford University School of Medicine, Stanford, California; <sup>3</sup>Department of Medical Microbiology and Immunology of Infection, Institute for Infection Medicine, Benjamin Franklin Medical Center, Free University of Berlin, Berlin, Germany

İlk trimesterda yüksek avidite sonucu, fetal infeksiyon olasılığını dışlayabilir mi?

Gebeliğin başlangıcında edinilmiş infeksiyona bağlı konjenital bulaş olasılığı sıfıra yakındır.

## Effect of Testing for IgG Avidity in the Diagnosis of *Toxoplasma gondii* Infection in Pregnant Women: Experience in a US Reference Laboratory

Oliver Liesenfeld,<sup>1,2,3</sup> Jose G. Montoya,<sup>1,2</sup>  
Sandra Kinney,<sup>1</sup> Cynthia Press,<sup>1</sup>  
and Jack S. Remington<sup>1,2</sup>

<sup>1</sup>Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto, and <sup>2</sup>Division of Infectious Diseases and Geographic Medicine, Department of Medicine, Stanford University School of Medicine, Stanford, California; <sup>3</sup>Department of Medical Microbiology and Immunology of Infection, Institute for Infection Medicine, Benjamin Franklin Medical Center, Free University of Berlin, Berlin, Germany

İlk trimesterda yüksek avidite sonucu;  
hasta izlemi ihtiyacını,  
amniyotik sıvıda PCR incelemesini,  
annede spiramisin tedavisini,  
hastanın anksiyetesini,  
maliyeti azaltır.

## Effect of Testing for IgG Avidity in the Diagnosis of *Toxoplasma gondii* Infection in Pregnant Women: Experience in a US Reference Laboratory

Oliver Liesenfeld,<sup>1,2,3</sup> Jose G. Montoya,<sup>1,2</sup>  
Sandra Kinney,<sup>1</sup> Cynthia Press,<sup>1</sup>  
and Jack S. Remington<sup>1,2</sup>

<sup>1</sup>Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto, and <sup>2</sup>Division of Infectious Diseases and Geographic Medicine, Department of Medicine, Stanford University School of Medicine, Stanford, California; <sup>3</sup>Department of Medical Microbiology and Immunology of Infection, Institute for Infection Medicine, Benjamin Franklin Medical Center, Free University of Berlin, Berlin, Germany

İlk trimesterda IgG avidite testi, yeni edinilmiş *T. gondii* infeksiyonunun dışlanması için değerli bir doğrulama testidir.

## VIDAS Test for Avidity of *Toxoplasma*-Specific Immunoglobulin G for Confirmatory Testing of Pregnant Women

Jose G. Montoya,<sup>1,2\*</sup> Oliver Liesenfeld,<sup>1†</sup> Sandra Kinney,<sup>1</sup> Cynthia Press,<sup>1</sup> and Jack S. Remington<sup>1,2</sup>

*Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto,<sup>1</sup> and  
Department of Medicine and Division of Infectious Diseases and Geographic Medicine, Stanford University  
School of Medicine, Stanford,<sup>2</sup> California*

VIDAS Toxo IgG avidite kiti ile yüksek avidite, son 4 ay içindeki infeksiyonu dışlar.



## **IgA Antibodies for Diagnosis of Acute Congenital and Acquired Toxoplasmosis**

**Pamela Stepick-Biek, Philippe Thulliez,  
Fausto G. Araujo, and Jack S. Remington**

*From the Department of Immunology and Infectious Diseases,  
Research Institute, Palo Alto Medical Foundation, and Division of  
Infectious Diseases, Department of Medicine, Stanford University  
School of Medicine, Stanford, California; and Laboratoire de Serologie  
Neonatale et de Recherche sur la Toxoplasmose,  
Institut de Puericulture de Paris, France*

An ELISA for IgA toxoplasma antibodies was positive in 12 pregnant women who seroconverted during gestation. Positive IgA titers were also noted in 10 individuals with biopsy-proven toxoplasmic lymphadenitis; the highest titers were noted in the first months following onset of clinical signs. Toxoplasma IgA antibodies were also demonstrable in 8 of 9 infants/fetuses with congenital toxoplasma infection. In some, IgM antibodies could not be demonstrated. Among 20 patients with AIDS and biopsy-proven toxoplasmic encephalitis, only 1 had IgA antibodies. None of 20 individuals with chronic toxoplasma infection had demonstrable IgA antibodies. Demonstration of IgA toxoplasma antibodies should be useful for diagnosis of recently acquired infection and for diagnosis of the infection in the fetus and newborn.

## *T. gondii* IgA antikorları

akut infeksiyonun gösterilmesinde,  
fetüs ve yenidoğanda tanıda önemli,  
duyarlılıkları IgM testlerinden daha iyi.

## Role of Specific Immunoglobulin E in Diagnosis of Acute Toxoplasma Infection and Toxoplasmosis

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### *T. gondii* IgE antikorları

Akut infeksiyon tanısında önemli,  
IgM ve IgA'dan daha kısa süre pozitif  
kalır.

Toksoplazmozun serolojik tanısında hiçbir test “mükemmel” değildir ve bu testler birlikte kullanılmalıdır.



## Olgu

32 yaşında,  
12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok,  
Herhangi bir yakınması yok,  
*T. gondii* IgM (+), IgG (+),  
IgG avidite düşük.

## VIDAS Test for Avidity of *Toxoplasma*-Specific Immunoglobulin G for Confirmatory Testing of Pregnant Women

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*Department of Immunology and Infectious Diseases, Research Institute, Palo Alto Medical Foundation, Palo Alto,<sup>1</sup> and  
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School of Medicine, Stanford,<sup>2</sup> California*

Kimi hastalarda düşük aviditeli antikörlerin aylarca, hatta bir yıldan uzun süre saptanabilmesi nedeniyle, avidite testi yeni edinilmiş infeksiyon tanısı için güvenilir değildir.

Avidite testi, düşük ya da sınırda sonuçların yanlış değerlendirilmesi olasılığı nedeniyle, IgM / IgG pozitif gebelerde “tek” doğrulama testi olarak kullanılmamalıdır.



Olgu

*T. gondii* IgM (+), IgG (+),  
IgG avidite düşük.

Tedavi

Fetal tanı



Amniyotik  
sıvı PCR



USG

Gebe, akut toksoplazmoz dışlanamıyor

<18 hafta

>18 hafta

Spiramisin  
Fetal USG  
Amniyotik sıvı  
PCR, >18. hafta

Primetamin+sülfadiazin  
+folinik asit  
Fetal USG  
Amniyotik sıvı PCR

PCR (-)  
USG normal

PCR (+)  
USG anormal

PCR (-)  
USG normal

Spiramisin  
devam

Primetamin+  
Sülfadiazin+  
Folinik asit

PSF devam  
ya da  
spiramisin

## PRENATAL DIAGNOSIS OF CONGENITAL TOXOPLASMOSIS WITH A POLYMERASE-CHAIN-REACTION TEST ON AMNIOTIC FLUID

PATRICK HOHLFELD, M.D., FERNAND DAFFOS, M.D., JEAN-MARC COSTA, PH.D., PHILIPPE THULLIEZ, M.D., FRANÇOIS FORESTIER, PH.D., AND MICHEL VIDAUD, PH.D.

**Abstract Background.** Congenital infection with *Toxoplasma gondii* can produce serious sequelae. However, there is little consensus about screening during pregnancy, and the tests used to establish a prenatal diagnosis of toxoplasmosis are complex and slow. We evaluated a simpler approach that is based on a polymerase-chain-reaction (PCR) test.

**Methods.** Prenatal diagnostic tests, including ultrasonography, amniocentesis, and fetal-blood sampling, were performed in 2632 women with *T. gondii* infection acquired during pregnancy. In 339 consecutive women, a competitive PCR test for *T. gondii* was performed on amniotic fluid, and its results were compared with those of conventional diagnostic tests. The PCR test targets the *B1* gene of *T. gondii*, uses an internal control, and can be completed in a day. Positive tests were confirmed by serologic testing

of newborns or by autopsy in terminated pregnancies.

**Results.** Overall, the risk of fetal infection was 7.4 percent, but it increased sharply with gestational age. Congenital infection was demonstrated in 34 of 339 fetuses by conventional methods, and the PCR test was positive in all 34. In three other fetuses, only the PCR test gave positive results, and follow-up testing confirmed the presence of congenital toxoplasmosis. The PCR test gave one false negative result but no false positive results. The PCR test performed better than conventional parasitologic methods (sensitivity, 97.4 percent vs. 89.5 percent; negative predictive value, 99.7 percent vs. 98.7 percent).

**Conclusions.** For the prenatal diagnosis of congenital *T. gondii* infection, an approach based on a PCR test performed on amniotic fluid is rapid, safe, and accurate. (N Engl J Med 1994;331:695-9.)

Amniyotik sıvıda *T. gondii* DNA PCR  
optimal zaman gebeliğin 18. haftası,  
akut infeksiyonun üzerinden en az 4  
hafta geçmeli...



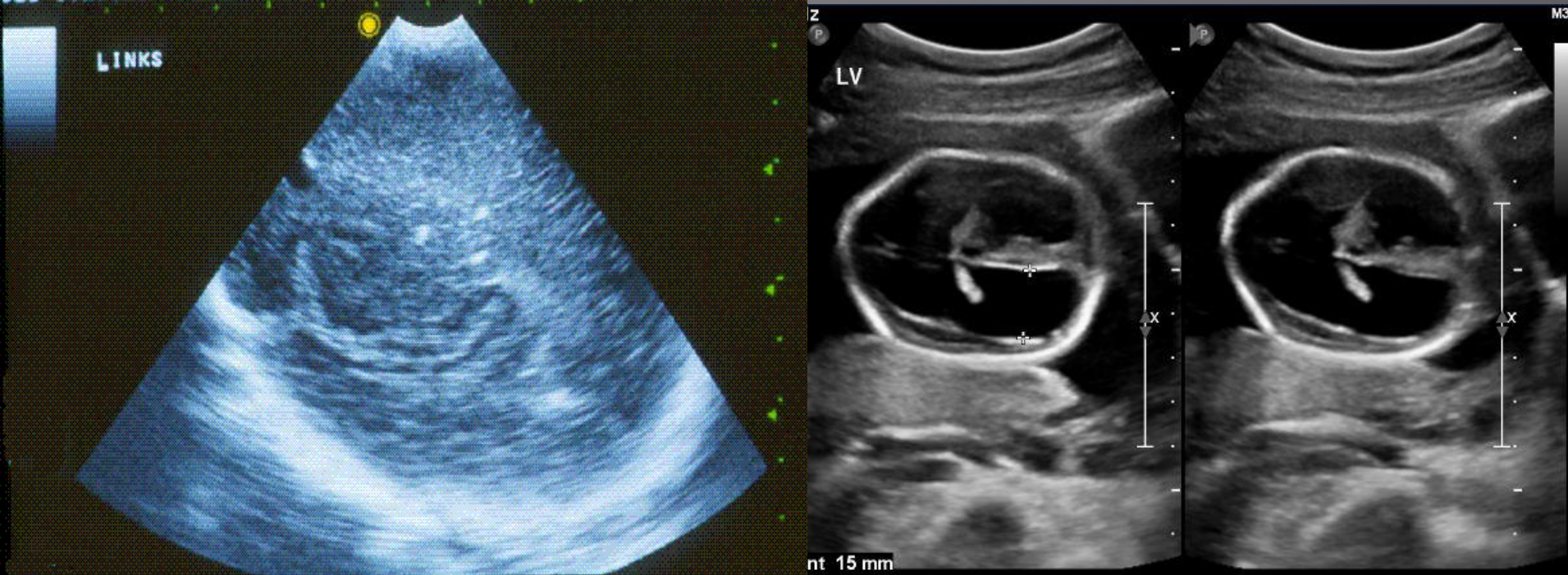
## Amniyotik sıvıda *T. gondii* DNA PCR

duyarlılık %64-92

özgüllük %100

npd %88-99

ppd %100



## USG

bulgular genellikle özgül değildir,  
intrakraniyal hiperekojen odak ya da  
kalsifikasyonlar,  
bilateral, simetrik ventriküler  
dilatasyon,  
intrahepatik dansiteler, asit,  
plasental kalınlaşma, hiperdensite

## Spiramisin

3x1 gr/gün,  
kimi çalışmalarda bulaşı %60 azalttığı  
bildirilmiş,  
plasentayı iyi geçmez, fetal  
infeksiyonun tedavisi için uygun değildir,  
amniyotik sıvı PCR (-) olsa dahi,  
doğuma dek kullanılmalıdır.

## Primetamin

en etkili antitoksoplazma ajan,  
folik asit antagonisti,  
doz ilişkili kemik iliği süpresyonu,  
teratojenik potansiyel,  
2x50 mg/gün 2 gün yükleme, 50  
mg/gün

## Sülfadiazin

primetamin ile sinerjistik etki,  
75 mg/kg yükleme, 2x50 mg/gün (max  
4 gr)

## Folinik asit

kemik iliği süpresyonunu önler,  
10-20 mg/gün

## Effect of timing and type of treatment on the risk of mother to child transmission of *Toxoplasma gondii*

### European Multicentre Study on Congenital Toxoplasmosis\*

*Participants are listed on page 119*

**Objective** To determine the effects on mother to child transmission of the timing and type of prenatal treatment, taking into account gestational age at maternal seroconversion.

**Design** Prospective cohort study.

**Setting** European centres offering prenatal screening for toxoplasmosis.

**Population** Children born to a cohort of pregnant women with toxoplasma infection.

**Methods** We determined the effects on mother to child transmission of the interval between seroconversion and start of treatment (treatment delay), and the type of treatment, taking into account gestational age at maternal seroconversion.

Prenatal tedavide kullanılan ilacın ya da tedaviye başlama zamanının anneden fetüse bulaş riskine etkisi yoktur.



# Effectiveness of prenatal treatment for congenital toxoplasmosis: a meta-analysis of individual patients' data

*The SYROCOT (Systematic Review on Congenital Toxoplasmosis) study group\**

## Summary

**Background** Despite three decades of prenatal screening for congenital toxoplasmosis in some European countries, uncertainty remains about the effectiveness of prenatal treatment.

*Lancet 2007; 369: 115–22*

\*Members listed at end of report

**Methods** We did a systematic review of cohort studies based on universal screening for congenital toxoplasmosis. We did a meta-analysis using individual patients' data to assess the effect of timing and type of prenatal treatment on mother-to-child transmission of infection and clinical manifestations before age 1 year. Analyses were adjusted for gestational age at maternal seroconversion and other covariates.

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**Findings** We included 26 cohorts in the review. In 1438 treated mothers identified by prenatal screening, we found weak evidence that treatment started within 3 weeks of seroconversion reduced mother-to-child transmission compared with treatment started after 8 or more weeks (adjusted odds ratio [OR] 0.48, 95% CI 0.28–0.80;  $p=0.05$ ). In 550 infected liveborn infants identified by prenatal or neonatal screening, we found no evidence that prenatal treatment significantly reduced the risk of clinical manifestations (adjusted OR for treated vs not treated 1.11, 95% CI 0.61–2.02). Increasing gestational age at seroconversion was strongly associated with increased risk of mother-to-child transmission (OR 1.15, 95% CI 1.12–1.17) and decreased risk of intracranial lesions (0.91, 0.87–0.95), but not with eye lesions (0.97, 0.93–1.00).

26 kohort çalışma,  
1745 infekte anne,  
1438 tedavi alan olgu





# Effectiveness of prenatal treatment for congenital toxoplasmosis: a meta-analysis of individual patients' data

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## Summary

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Serokonversiyondan sonra 3 hafta içinde başlanan tedavi, bulaşı azaltmaktadır.

Gebelik sırasında tedavi edilen annelerin bebeklerinde ciddi nörolojik sekel ve postnatal ölümdede azalma vardır.

Mevcut alıřmaların hibiri klinik fayda olasılıđını tamamen dıřlamaz,  
Özellikle yeni alıřmalarda erken tedavinin yararı gösterilmiřtir,

Akut *T. gondii* infeksiyonu olasılıđı yüksek olan gebelerde spiramisin ya da primetamin+sülfadiazin tedavisi önerilmeye devam edilmektedir.





## Olgu

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12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok,  
Herhangi bir yakınması yok,  
*T. gondii* IgM (+), IgG (+),  
IgG avidite düşük.

Spiramisin 3x1 gr/gün, doğuma dek,  
18. haftada amniyosentez, PCR (-),  
Fetal USG normal.



## Olgu

32 yaşında,  
12 haftalık gebe,  
Primigravid,  
Antenatal izleminde özellik yok,  
Herhangi bir yakınması yok,  
*T. gondii* IgM (+), IgG (+),  
IgG avidite düşük.

Sonuç olarak,  
gebelerde IgM pozitifliği her zaman yeni edinilmiş infeksiyon anlamına gelmez,  
düşük IgG aviditesi her zaman yeni edinilmiş infeksiyon anlamına gelmez,  
18. haftadan sonra amniosentez ve *T. gondii* DNA PCR ile USG izlemi önemlidir,  
yeni edinilmiş infeksiyonun dışlanamadığı gebelerde tedavi planlanmalıdır.