Molecular diagnosis of toxoplasmosis at the onset of symptomatic primary infection: A straightforward alternative to serological examinations

Maude Lévêque, Delphine Chiffre, Chloé Galtier, Sahar Albaba, Christophe Ravel, Laurence Lachaud, Anaïs Iemmi, Pierre Flori, Yvon Sterkers

To cite this version:

Maude Lévêque, Delphine Chiffre, Chloé Galtier, Sahar Albaba, Christophe Ravel, et al.. Molecular diagnosis of toxoplasmosis at the onset of symptomatic primary infection: A straightforward alternative to serological examinations. International Journal of Infectious Diseases, Elsevier, 2019, 79, pp.131-133. 10.1016/j.ijid.2018.11.368. hal-02408741

HAL Id: hal-02408741
https://hal.umontpellier.fr/hal-02408741
Submitted on 13 Dec 2019

HAL is a multi-disciplinary open access archive for the deposit and dissemination of scientific research documents, whether they are published or not. The documents may come from teaching and research institutions in France or abroad, or from public or private research centers.

L’archive ouverte pluridisciplinaire HAL, est destinée au dépôt et à la diffusion de documents scientifiques de niveau recherche, publiés ou non, émanant des établissements d’enseignement et de recherche français ou étrangers, des laboratoires publics ou privés.

Distributed under a Creative Commons Attribution - NonCommercial - NoDerivatives| 4.0 International License
Case Report

Molecular diagnosis of toxoplasmosis at the onset of symptomatic primary infection: A straightforward alternative to serological examinations

Maude F. Lévêque, Delphine Chiffre, Chloé Galtier, Sahar Al Baba, Christophe Ravel, Laurence Lachaude, Anaïs Lemi, Pierre Flori, Yvon Sterkers

Article history:
Received 13 November 2018
Received in revised form 27 November 2018
Accepted 28 November 2018

Corresponding Editor: Eskild Petersen, Aarhus, Denmark

Keywords:
Acute toxoplasmosis
Myopericarditis
Immunocompetent
Toxoplasma gondii
PCR
Second serological control
Molecular diagnosis

Abstract

Myopericarditis is a rare but well-documented clinical presentation of primary Toxoplasma gondii infection in immunocompetent patients. Here, early detection of Toxoplasma DNA in the peripheral blood by PCR allowed the diagnosis of acute toxoplasmosis while serological tests were negative. Additional serological evaluations 2 weeks later confirmed the diagnosis and showed that cardiac manifestations occurred before seroconversion. This highlights the importance of a second serological control in the case of a suspected active infection. Overall, we show here that PCR testing for Toxoplasma is a sensitive and straightforward alternative to serological examinations.

Introduction

Toxoplasma gondii is the causative agent of toxoplasmosis and one of the most prevalent zoonotic agents. It infects approximately one-third of the human population globally (Montoya and Liesenfeld, 2004). This parasite spans distinct biological barriers during acute infection and eventually persists as tissue cysts to escape the host immune system. Humans are generally infected through the consumption of undercooked meat or vegetables and water contaminated by cat faeces.

Uncontrolled parasite replication during primary infection in the foetus and reactivation of tissue cysts in the immunocompromised host can lead to life-threatening involvement of one or more organs. Myocarditis is a well-known complication of toxoplasmosis, most often associated with HIV infection (Kirchoff et al., 2004). Cardiac clinical manifestations result from active parasite invasion of the pericardium and myocardium, producing local inflammatory responses and potential subsequent cardiomyopathies (Nunes et al., 2017).

This article reports the diagnosis of Toxoplasma myopericarditis, occurring before seroconversion for T. gondii, by detection of parasite DNA in the peripheral blood of a patient with no laboratory or clinical evidence of immunosuppression. The diagnosis of acute toxoplasmosis in immunocompetent subjects is discussed.

Case report

A 23-year-old man was admitted to the Emergency Room of the Academic Hospital of Montpellier on March 13, 2018. He complained of intense chest pain not relieved by paracetamol, which had started 2 days ago. At the time of admission, a physical examination was normal and the patient appeared febrile, with normal blood pressure and a rhythmic heart beat of 87 bpm. The pain was typical of pericarditis: it was retrosternal, constrictive,

* Corresponding author at: Research Unit “MIPEGEC” (CNRS 5290/IRD 224/UM), University of Montpellier, Montpellier, France.
E-mail address: Yvon.sterkers@umontpellier.fr (Y. Sterkers).

https://doi.org/10.1016/j.ijid.2018.11.368
1201-9712/© 2018 The Author(s). Published by Elsevier Ltd on behalf of International Society for Infectious Diseases. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).
increased with deep inspiration, and was relieved by anteflexion and after treatment with acetylsalicylic acid 1g. There was no evidence of pericardial friction, which is usually tickle, fleeting, and does not eliminate the diagnosis if absent. An electrocardiogram and echocardiogram showed no particular abnormality. Troponin, a marker of myocardial necrosis, was elevated at 31.3 ng/l (reference value <14 ng/l) as was C-reactive protein (CRP) at 76.1 ng/l (reference value <5 mg/l), suggesting an infectious myopericarditis in the absence of an alternative aetiology.

The patient was transferred to the cardiology intensive care unit the day after admission. Serological and molecular testing was negative for herpes simplex virus 1 and 2, hepatitis B and C virus, HIV, and human herpes virus 6 and 8, and consistent with past exposure to rubella virus, cytomegalovirus, parvovirus B19, and Epstein–Barr virus. The results of immunological analyses were normal (lymphocytes, plasma immunoelectrophoresis, autanti-bodies).

Three days after the appearance of the symptoms, serological tests for toxoplasmosis were negative; however, PCR to detect T. gondii DNA in the peripheral blood was positive (Table 1). Recent primary infection by the parasite was therefore considered the aetiological factor of the myopericarditis. The patient did not present any symptoms of cervical lymphadenopathy or myalgia, which may occur during the initial stages of T. gondii infection (Montoya and Liesenfeld, 2004). However, he complained of intermittent abdominal pain a few days before his admission to the emergency room.

During his hospitalization, the patient’s clinical condition improved and levels of troponin and CRP normalized 3 days after the appearance of the cardiac symptoms. Treatment based on acetylsalicylic acid 1g three times a day for 1 month, lasoprazole 15mg once a day for 1 month, colchicine 1mg once a day for 3 months, and bisoprolol 1.25mg once a day for 3 months was prescribed. The patient did not receive any drugs for toxoplasmosis and was discharged 4 days after admission with an appointment for cardiac magnetic resonance imaging. This revealed probable lateral myocarditis. To confirm the diagnosis of toxoplasmosis, biological follow-up was requested. Two weeks after the pericardi-tis episode with elevation of troponin levels, PCR no longer detected T. gondii DNA in the peripheral blood (Table 1), suggesting that the circulation of parasites had been very transient. However, IgG and IgM were found positive, with a consistent low antigen affinity of IgG at 0.14 index (high >0.6), confirming a recent Toxoplasma seroconversion (Table 1).

Discussion

Toxoplasma primary infection is asymptomatic and harmless for the vast majority of healthy individuals, causing mononucleosis-like symptoms with cervical lymphadenopathy in 10% of subjects (Montoya and Liesenfeld, 2004). Primary infections with atypical strains from South America can be associated with severe clinical involvement in the immunocompetent host (Sobanski et al., 2013).

In the case presented here, the patient’s history revealed that he had eaten raw horsemeat at a restaurant less than 2 weeks before the appearance of his cardiac symptoms, while no other risk factor was found. The meat served at the restaurant during this period of time had come from Europe and was, therefore, unlikely contaminated by atypical strains. Despite the attempt to genotype the Toxoplasma DNA isolated from the patient’s blood, it was not possible to confirm this assumption due to the low parasite load.

Toxoplasma pericarditis and myocarditis have already been documented in immunocompetent patients (Bousquet et al., 2016; Chandenier et al., 2000; Montoya et al., 1997; Mroczek-Czernecka et al., 2006; Paspalaki et al., 2001; Pergola et al., 2010; Roubille et al., 2012; Sano et al., 2000), although the number of cases is low. In these previous cases, serological testing was generally used to link the cardiac complications to toxoplasmosis. Indeed, guidelines for the diagnosis of acute toxoplasmosis in immunocompetent patients state the requirement for two serological tests separated by 3 weeks. However, as illustrated here, a number of symptomatic primary infections may be undiagnosed at the first serological examination, due to the delayed synthesis of specific antibodies.

This case highlights the importance of performing blood PCR analysis, which is likely to be the first positive biological test, soon after the appearance of the cardiac symptoms. Indeed, serological conversion for toxoplasmosis was evidenced after the patient’s recovery from his cardiac disorders (i.e., relief of chest pain and normalization of troponin levels). Therefore, in the absence of molecular diagnosis and the second serological control, the patient’s transient disorders would not have been associated with active toxoplasmosis. The identification of Toxoplasma as the aetiological factor of myopericarditis is, however, necessary in order to exclude other probably more severe diagnoses.

In conclusion, when symptomatic Toxoplasma primary infection is suspected, physicians should consider blood PCR as a sensitive and straightforward approach for diagnosis.

Acknowledgements

We are grateful to Professor Marie-Laure Dardé and the Biological Reference Centre for Toxoplasmosis, Limoges, for genotyping attempts. We thank Florence Michel and Sylvie Douzou for their technical assistance.

Funding sources

We acknowledge the financial support of the French Ministry of Health (Santé Publique France) and “RSI Assurance Maladie Professions Libérales – Provinces C.A.M.P.L.P.” for buying the LightCycler 480 real-time PCR equipment (Roche). The study sponsors had no role in the study design, the collection, analysis,
and interpretation of the data, the writing of the manuscript, or in the decision to submit the manuscript for publication.

**Ethical approval**

We read and complied with the policy of the journal on ethical consent. The study was conducted in accordance with the regulations of the local medical ethics committee of the Hospital University Centre (CHU) of Montpellier, France, in line with the revised Declaration of Helsinki.

**Conflict of interest**

The authors declare no conflict of interest.

**Author contributions**

Study design: MFL and YS; data collection: MFL, DC, CG, SA, AI, PF, YS; data analysis: MFL, DC, PF, YS; writing: MFL, DC, CR, LL, PF, YS.

**References**


