



# The mysteries of rapidly destructive arthrosis of the hip joint: a systemic literature review

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**Abstract:** Rapidly destructive arthrosis (RDA) is considered a rare and poorly diagnosed disease but is now seen more frequently in practice due to ageing populations. The most typical radiological features are flat femoral heads, absence of articular cartilage, subchondral bone destruction and signs of joint effusion. These features could be found on X-ray or magnetic resonance imaging (MRI). Surgeons should consider the presence of RDA when patients show rapid femoral head destruction. The purpose of this study is to review the distinct clinical features and successful treatments which may lead to the diagnosis and early handling of RDA. A comprehensive review of the literature was undertaken using Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines with no language restrictions. Overall 23 publications with 17 detailed cases of RDA met the inclusion criteria. We found that the only prevalent factors associated with RDA were: (I) age greater than 60 years; (II) female gender; (III) presence of underlying systemic diseases such as rheumatoid arthritis, diabetes mellitus or systemic lupus erythematosus. Further studies should be conducted to clarify the histopathology and define the diagnosis as well as the treatment.

**Keywords:** Rapidly destructive arthrosis (RDA); risk factor; total hip arthroplasty; osteoarthritis (OA); systemic review

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## Introduction

Rapidly destructive arthrosis (RDA) is an unusual, poorly diagnosed disease. It was firstly reported by Forestier in 1957 and a standardized definition was provided by Lequesne in 1970: 50% articular space narrowing in 1 year or femoral head destruction >2 mm in 1 year (1). The disease diagnoses that are only based on radiographic

findings can be inconclusive. In some cases, they are easily misdiagnosed as rheumatoid arthritis, neuroarthropathy, septic arthritis, osteonecrosis of the femoral head, seronegative arthritis or osteoarthritis (OA). But in fact, the clinical, laboratory and pathologic symptoms of RDA are not in accordance with these diseases. The purpose of this systematic review is to investigate the clinical and

pathophysiology features to facilitate correct diagnosis and prompt appropriate treatment of RDA.

## Materials and methods

A comprehensive review of literature was undertaken using Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines with no language restrictions. Searches were conducted using the following databases: PubMed, Embase, Cochrane Library, Springer, and the Google Scholar search tool. The following keywords were used: “rapidly destructive osteoarthritis”, “rapidly destructive arthropathy”, “rapidly destructive coxarthrosis”, “rapidly destructive arthrosis”, “rapidly destructive arthrosis of the hip”, “rapidly progressive osteoarthritis”, “Postel’s osteoarthritis”, “destructive osteoarthritis” and “rapid destructive coxopathy”. Two researchers selected potentially relevant abstracts and obtained full copies of the articles. In addition, the references of the articles were reviewed.

### Criteria for eligibility

The studies selected were original clinical articles that addressed RDA in elderly patients with no language restrictions. Cases with pigmented villonodular synovitis, pathological bone fractures and neuroarthropathy—i.e., Charcot joint—were excluded. Date limits were set from the inception of the journal to March 2018.

### Data extraction

The following data were extracted from the eligible articles and case studies: type of study, age, gender, comorbidities, symptoms and signs, diagnostic modalities, treatment and outcome.

### Statistical analysis

As the majority of the data collected were from case reports and case series, statistical analysis was not possible. Descriptive statistics were employed where suitable.

## Results

### Literature search

After omitting repetitions and studies which did not fulfil the selection criteria, 23 case reports or case-control studies

were included in the analysis. All of these studies were retrospective. Fourteen were reported in Asia (2-15), six in Europe (16-21) and three in America (22-24). In total, 17 detailed patient cases were included (one article reported two cases, see *Table 1*). Seven case-control studies and case series including 164 patients were also studied (*Table 2*). We also reviewed additional relevant articles to facilitate the development of the discussion. The literature search flowchart could be found in *Figure 1*.

### Epidemiology

#### Age

A total of 17 cases reported the age of patients. The mean age was 69.2 years (range, 37–81 years). There were 3 patients under 60 years (17.6%) with a mean age of 50.3 years (range, 37–57 years), and 14 patients over 60 years (82.4%) with a mean age of 73.3 years (range, 66–81 years). The seven cohort studies reported a total of 164 cases, with a mean age of 70.5 years.

#### Gender

All studies (181 cases) included 37 males (20.4%) and 144 females (79.6%)

#### Disease bilaterality

Among all reported studies (181 cases), bilateral arthropathy was found in three cases (1.7%). In the majority (178 cases), RDA occurs unilaterally.

#### Diagnosis

The most commonly accepted diagnosis criteria of RDA was joint space loss occurring at a rate greater than 2 mm per year or if more than 50% of joint space was lost in one year, reported by Lequesne in 1970. RDA was often misdiagnosed as another disease, including rheumatoid arthritis, neuroarthropathy, septic arthritis, osteonecrosis of the femoral head, seronegative arthritis and osteoarthritis.

### Radiological findings

#### X-ray

In all patients, the weight-bearing area of the femoral head was flat. The articular cartilage was absent and the subchondral bone was destroyed.

#### Magnetic resonance imaging (MRI)

All patients had hip joint effusion and evidence of synovitis.

**Table 1** Clinical characteristics of the 17 patients

Author	Year	Age	Gender	BMI	Affected side	Comorbidities	Trauma	Course of disease	Blood test	Bone marrow edema	Diagnostic modalities	Early subchondral fracture
Huerfano	2017	76	F	–	Right	DM	No	6 W	Normal	Acetabulum, femoral head and neck and the irregular focal high-intensity bands	X-ray + MRI	Yes
Yamamoto	2010	57	F	19.7	Bilateral	Hyperlipidemia	No	10 M	Normal	–	X-ray	Yes
Celik	2015	78	M	–	Bilateral		–	2 Y	ESR =34 mm/h	–	X-ray	Yes
Laroche	2002	67	M	–	Left	Coronary heart disease diabetes mellitus occlusive arterial disease	–	3 M	ESR =25 mm/h; CRP =10 mg/L	Subchondral bone edema	X-ray + MRI	Yes
Yun	2012	67	F	19.9	Left	Rheumatoid arthritis	No	6 M	ESR =23 mm/h; CRP =79.5 mg/L; RF =50 IU/mL	–	X-ray	–
Yun	2012	67	M	16.9	Left	Rheumatoid arthritis	No	6 M	ESR =75 mm/h; CRP =418.5 mg/L; RAF =47 IU/mL	Severe cartilage erosion	X-ray + MRI	–
Yang	2011	66	M	–	Right	No	No	6 M	ESR =31 mm/h, CRP =24.5 mg/L	–	X-ray + MRI	Yes
Nishida	2005	74	F	22.8	Right	A history of a vertebral fracture	No	17 M	–	Femoral head	X-ray + MRI	YES
Fukui	2015	77	F	22.5	Left	No	No	5 M	–	Femoral head and the acetabulum	X-ray + MRI	Yes
Suzuki	2018	80	F	–	Right	Hypertension and dyslipidemia	No	3 M	–	–	X-ray + CT	Yes
Homma	2014	80	F	–	Left	Right postoperative THA	No	2 M	–	Acetabulum, femoral head and neck and the irregular focal high-intensity bands	X-ray + MRI	–

**Table 1** (continued)

Table 1 (continued)

Author	Year	Age	Gender	BMI	Affected side	Comorbidities	Trauma	Course of disease	Blood test	Bone marrow edema	Diagnostic modalities	Early subchondral fracture
Watarai	2008	81	F	22	Left	None	No	3 M	Normal	Acetabulum, femoral head and neck and the irregular focal high-intensity bands	X-ray + MRI	Yes
Lee	2015	37	F	22.3	Right	Systemic lupus erythematosus (SLE)	No	4–13 M	WBC =4,000/ $\mu$ L, PLT = $180 \times 10^3$ / $\mu$ L, C3 =90 mg/dL, C4 =20 mg/dL	Femoral head and the acetabulum	X-ray + MRI	Yes
Shu	2014	57	F	–	Right	RA	No	5 M	ESR =56 mm/h; CRP =106.4 mg/L; ANA =1:80	–	X-ray + MRI + CT	–
Marley	2013	74	F	–	Left	Osteoarthritis	No	9 M	Normal	–	X-ray	–
Hamada	2014	69	F	–	Left	Ochronosis	No	–	CRP =7.7 mg/L, HLA-B27(–)	Femoral head and the acetabulum	X-ray + MRI	Yes
Hama	2015	70	F	–	Right	Severe platelet deficiency, liver cirrhosis, immune thrombocytopenic purpura	No	3 M	CRP =27.7 / mg/L, RBC = $243 \times 10^4$ / $\mu$ L, PLT = $2.2 \times 10^4$ / $\mu$ L	–	X-ray	–

(–), absence.

Diffuse signal abnormalities were found in the marrow of the femoral head and neck. None of the patients exhibited chondrocalcinosis (pelvis, knees), osteoarthritis radiographically or histopathologically. Low signal intensity occupied the whole femoral head on T1-weighted images. The same area showed high signal intensity on T2-weighted images. A small low-intensity line was observed at the weight-bearing area (25).

#### Potential risk factors

##### Body mass index (BMI)

Seven cases reported the BMI of the patients. Six patients had the standard BMI (range, 19.7–22.8) and one had a

lower BMI (16.9).

##### Trauma

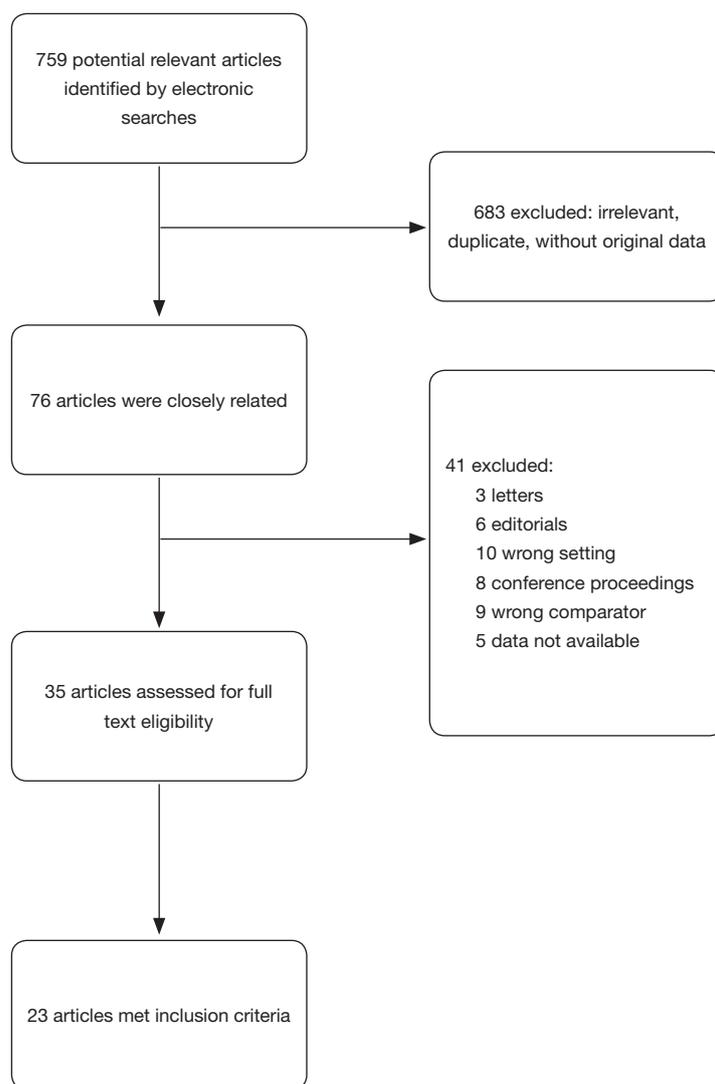
None of the cases reviewed showed evidence of trauma at the onset of disease.

##### Corticosteroid and alcohol

Only four patients had a history of glucocorticoid administration and two patients reported daily consumption of alcohol. One of them had both corticosteroid and alcoholism.

##### Secondary contralateral hip osteoarthritis

There was insufficient evidence to suggest that patients with



**Figure 1** Flow diagram of scientific literature search and study selection.

RDA have a higher risk of developing osteoarthritis in the contralateral hip than patients with OA (26).

#### **Bone mineral density (BMD)**

One study reported that no significant differences were observed in BMD between RDA and OA patients (27).

#### **Clinical characteristics**

##### **Symptoms and signs**

A total of 17 cases included the pain and limitations reported by patients suffering from RDA. Because of the intractable severity of pain, two patients underwent a total

hip replacement (2,9). Twelve patients were reported to have  $38.3 \pm 3.07$  (range, 33–42) points in the Harris Hip Score (15). Two patients were unable to walk because of the pain (8,11). One case reported the pain level as 9/10 by visual analog scale and their Barthel Index was 25 suggesting extreme dependence for others when carrying out daily activities.

##### **Comorbidities**

The cases reported hyperlipidemia, tonsillectomy, coronary heart disease, diabetes mellitus, occlusive arterial disease, rheumatoid arthritis, hypertension, systemic lupus erythematosus, ochronosis, severe platelet deficiency,

**Table 2** Summary of findings in the case series and cohort studies of RDA

Author	Year	Country	Sample size	Side	Gender	Control	Summary of findings
Yamamoto	2000	Japan	11	8 R/3 L	9 F/2 M	–	Subchondral insufficiency fracture resulting from osteopenia may lead to a rapid breakdown of the hip joint
Charrois	2001	France	100	–	86 F/14 M	100 (53 F/47 M)	The blood loss after THA performed for RDA is greater than for regular coxarthrosis
Boutry	2002	France	12	5 R/7 L	8 F/4 M	–	MR imaging can be valuable in the evaluation of such disorders
Yamakawa	2005	Japan	6	–	–	6	Hypervascularity of the granulation in the femoral head may be associated with bone and joint destruction
Ogawa	2007	Japan	10	–	10 F	40 F	The underlying mechanism of RDA is therefore associated with osteoclastogenesis in the synovium
Song	2015	Korea	19	–	6 F/13 M	40 (13 F/27 M)	Total perioperative blood loss was significantly greater in RDA than in ONFH in THA
Yuasa	2016	Japan	12	–	12 F	12 F	Cemented or cementless THA achieved a good midterm outcome.

liver cirrhosis, immune thrombocytopenic purpura, and esophageal varices as comorbidities of RDA.

### Medications

Two patients used hydroxychloroquine and methotrexate for rheumatoid arthritis. One used thrombopoietin receptor agonist and Kenketsu Glovenin-I for immune thrombocytopenic purpura (9).

### Blood tests

Nine cases reported normal blood tests in patients while seven cases reported higher than average erythrocyte sedimentation rate (ESR) and C-reactive protein (CRP).

### Pathophysiology

#### Subchondral fracture and bone marrow edema

Of the cases with reported or clear radiographs, the subchondral fracture could be seen in 11 cases. There were nine cases reporting bone marrow edema in the acetabulum, femoral head and neck, or the irregular focal high-intensity bands.

#### Osteoclasts infiltration

Mature and activated osteoclasts were present in the

synovium of RDA patients. Tartrate resistant acid phosphatase (TRAP) positive multinuclear giant cells present in synovial membrane showed the nature of osteoclasts (12,13).

### Experience in treatment

Most patients were treated with nonsteroidal anti-inflammatory drugs (NSAIDs) and some patients were treated with alendronate sodium hydrate and alfacalcidol. They all reported negligible effects on the advance of the disease. Fifteen patients received total hip arthroplasty with no complications reported. Two other patients refused to have surgical treatment. The cohort studies also reported total hip arthroplasties (THAs) in 164 patients. The mean interval of time from hip pain onset to the surgery, reported in 16 cases, was 26.5 weeks, with a range between 6 and 54 weeks. The total blood loss was significantly greater in arthroplasty performed for RDA than patients with osteoarthritis. Sex, age and extra body weight index (BMI) had no significant effect on bleeding (14-16). The mean Harris Hip Score improved significantly after surgery (15). THA achieved a good midterm outcome comparable to that for patients with primary or secondary osteoarthritis (15,28). Complications of the surgery were not different

from complications suffered by osteoarthritis patients.

### *Other findings*

The number of TRAP-positive multinuclear giant cells present in the synovial membrane obtained from RDA patients was significantly larger than that obtained from OA patients (13). The osteoclast count on the bone surface from the RDA patients was greater than that from the osteoarthritic patients (12).

### **Discussion**

RDA is an unusual, poorly diagnosed disease whose pathophysiology and etiology are still unknown. Various aspects of RDA continue to attract a lot of attention from both researchers and clinicians. In this literature review, we searched potentially relevant articles in the databases and reviewed the references of the articles. We then summarized the epidemiology and clinical characteristics of RDA from all the cases reported so far, and investigated the potential risk factors, diagnostic modalities, and experience in treatment. Accordingly, this article aims to provide evidence for improving the diagnosis and treatment of RDA.

Several potential risk factors have been proposed in the literature, including: (I) ageing; (II) female gender; (III) underlying health problems such as rheumatoid arthritis, diabetes mellitus and systemic lupus erythematosus; (IV) lower bone mineral density; (V) higher BMI; (VI) medicinal drug use including NSAIDs and corticosteroid; (VII) alcoholism (2-17,19-24,27,29). In this study, we evaluated the risk factors and found the only prevalent factors were: (I) age greater than 60 years; (II) female gender; (III) underlying systemic disease such as rheumatoid arthritis, diabetes mellitus or systemic lupus erythematosus (2-17,19-21,23,24). There is insufficient evidence to support the relationship between BMD, BMI, medication and alcoholism and the occurrence of RDA (2,3,5,8,10,23,27,29). We did not find these factors to increase the risk of RDA.

Most patients consulted a physician because of hip pain that had appeared without previous trauma or falls, and most of the disease progression occurred within one year. As a result, these changes were described as “rapid” and “destructive” in literature. Many of the patients were able to walk during their initial medical consultations, which may lead to the doctor failing to make an accurate diagnosis. Some studies mentioned that RDA was caused by simultaneous bilateral shoulder joint collapse within

a very short time, with minimal or low mechanical stress and severe osteoporosis (30). Although disease diagnosis is possible through physical examination and medical history, the first symptoms of RDA are neither specific to the disease nor clear. We suggest that any patient who has the above-mentioned high-risk factors should suggest the possibility of RDA to their medical practitioner and that further diagnostic measures should be taken.

Blood tests, including CRP and ESR, are nonspecific in this disease. We recommend an X-ray of the hip as the initial radiological examination; doctors were able to make an RDA diagnosis in all of the reported cases that had X-rays. In all patients, the X-rays showed that the weight-bearing area of the femoral head was flattened. Joint space loss at a rate greater than 2 mm per year, or if more than 50% of joint space had been lost in 1 year, were considered as RDA (1,31,32). MRI was also used in many cases and significant changes were found. It was reported that a unique presentation of severe bone destruction as a manifestation of chronic myeloid leukemia in the absence of blast crisis (33). There were bone marrow edema on acetabulum, femoral head and neck and the weight-bearing area and the articular cartilage was absent with the subchondral insufficiency fracture or subchondral bone destroyed (2,3,5-8,10,34,35).

Physicians should be aware of antidiastoles when making a diagnosis of RDA. Gorham-Stout Syndrome (Gorham's massive osteolysis) and RDA of the hip show similarities in the idiopathic rapid disappearance of bones. However, histological examinations have revealed that Gorham's massive osteolysis is associated with angiomatosis of blood vessels and sometimes with lymphatics (12). Charcot's joint (neuropathic osteoarthropathy) and RDA have similarities in the slight clinical signs with serious imaging manifestation. Diabetes mellitus is the leading cause of neuropathic osteoarthropathy, with additional associations including syringomyelia, meningocele and multiple sclerosis (36). It is easy to mistake RDA symptoms for symptoms of rheumatoid arthritis, septic arthritis, osteonecrosis of the femoral head, seronegative arthritis or osteoarthritis. The diagnosis of RDA should be exclusive.

We recommend direct joint replacement when the diagnosis is confirmed. Although cases with delayed replacement surgery showed no signs of malignancy or contralateral involvement (26), these patients showed negligible results on conservative treatment. Although Low-dose aspirin and monoclonal antibody against human receptor activator of nuclear factor- $\kappa$  B ligand (RANKL) were benefit for maintaining bone mass and qualities by

activation of osteoblastic bone formation and inhibition of osteoclast activities via cyclooxygenase-independent manner (37,38), the use of drugs such as NSAIDs, alendronate sodium hydrate and alfacalcidol had only mild or even no effect on the patients with RDA (3,5,6,20,24,39,40). Total hip replacement is probably the treatment of choice for these patients. In 100 RDA patients treated with cemented THA and followed up for a mean of 7.83 years, the Merle d'Aubigne score was excellent or very good in 95 hips (41). Yuasa *et al.* reported that after an average follow-up duration of 9.3 years, the mean Harris Hip Score improved from 38.3 to 81.1 in RDA patients and from 43.6 to 84.2 in conventional osteoarthritis controls (15). During surgery, the total blood loss of patients with hip osteoarthritis was significantly less than patients with RDA (9,14,16,35). Sex, age and extra body weight (BMI) had no significant effect on bleeding (14-16). The present study cannot explain the hemorrhagic nature of THA performed for RDA. We suppose that the excessive bleeding may be a result of the tissue edema or bone surface bleeding due to subchondral fractures. Operative time and the complexity of reconstructive efforts could be significantly reduced if the correct diagnosis was made early and surgical procedure was taken in time (32). Most RDA patients had the disease in a single joint and a few patients were reported as having bilateral RDA (21,23). Patients with RDA may not have a higher risk of developing osteoarthritis in the contralateral hip after THA, than patients with osteoarthritis (26). But chronic lameness could lead to spinal diseases such as lumbar scoliosis. Although outcomes in the reviewed cases appeared to be satisfactory, this cannot be confirmed because of the small number of cases with follow-up.

Bone is in a dynamic process of continuous remodeling which helps to regulate calcium homeostasis, repair micro-damage to bones from everyday stress, and to shape the skeleton during growth (42). Although the rapid clinical course and bone destruction in patients with RDA differ from those generally seen with a degenerative process, previously published reports have suggested that this disease is degenerative in nature and is a form or subset of OA (12,13,38). The reason why some joints undergo rapid and progressive destruction remains unclear. If fragility is the only cause of subchondral fracture in RDA, then fractures might occur at the femoral neck rather than at the head beneath the cartilage (43). Many paracrine or autocrine factors that stimulate osteoclastic differentiation and function in the active area have been studied (44,45). In patients with progressively destructive arthropathy,

it has been reported that osteoclasts with vascular-rich granulation could be detected (12,38,46). The osteoclast count was significantly greater in rapidly destructive arthropathic patients than in osteoarthritic patients (12,38) and an increase in osteoclasts was found in the active areas of the hip in patients with RDA (12,47). Several reports have shown that the elevated levels of bone-resorptive enzymes, such as interleukin-6 (IL-6), IL-1 $\beta$ , and matrix metalloproteinases, were associated with RDA of the hip (12). Hypervascularity was also found to be an important factor associated with the destruction of the bone and joint in RDA patients. Endothelium could play an important role in the recruitment of osteoclast precursors to sites of bone resorption because osteoclast precursors could adhere to the endothelium and be activated with the IL-6 secreted by endothelial cells (12,13,46).

The strengths of this study include a comprehensive search and analysis of world literature with no year or language limitations, and the relatively complete information provided in the reviewed cases. The main limitation is the small number of cases previously reported. Further studies on this disease might improve our knowledge of the management of RDA.

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