

Fever of Unknown Origin*	
Definition	Fever without a source for >7-10 days
Differential	<ul style="list-style-type: none"> • Bacterial: endocarditis, mastoiditis, sub-diaphragmatic abscess, liver abscess, perinephric abscess, pyelonephritis, pelvic abscess, osteomyelitis, TB, salmonellosis (including typhoid), lymphogranuloma venereum, brucellosis, cat-scratch disease, leptospirosis, tularemia, psittacosis, tick-borne disease (e.g. Anaplasma, Babesia), Q fever, RMSF • Viral: adenovirus, arboviruses (e.g. West Nile, dengue), primary HIV, CMV, EBV, HBV, HCV • Fungal: blastomycosis, histoplasmosis • Parasitic: malaria, toxoplasmosis, visceral larva migrans • Granulomatous: sarcoidosis, granulomatous colitis • Collagen Vascular Disease: systemic juvenile idiopathic arthritis, polyarteritis nodosa, SLE • Malignancy: leukemia, lymphoma, neuroblastoma, Langerhans cell histiocytosis • Miscellaneous: diabetes insipidus, drug fever, Kawasaki disease, familial dysautonomia (Riley-Day Syndrome), familial Mediterranean fever or other periodic fever syndromes, HLH, infantile cortical hyperostosis (Caffey Syndrome), pancreatitis, serum sickness, ulcerative colitis, thyrotoxicosis
Workup	<ul style="list-style-type: none"> • History with ROS, travel history, animal exposures, outdoor activities, insect bites, food exposures, sexual history, IV drug use • Exam: skin exam, LN palpation, joint exam • Labs: CBC with differential, UA/UCx, BCx, HIV, LFTs, LDH, CPK, ESR/CRP, ANA, TST/IGRA, LDH/Uric acid • Imaging: CXR to start; may require abdominal axial imaging (MRI vs. CT) • Additional work-up as indicated by history and physical and decided upon with guidance from consulting teams and radiology
Treatment	<ul style="list-style-type: none"> • Unless patient is very ill, empiric antimicrobial therapy should be avoided as it often delays diagnosis • Can observe fever pattern for diagnostic purposes before treating fever • Glucocorticoids or other immunosuppressive therapy should be withheld until infectious etiology is adequately ruled out