## Looking for Trouble: The Work up of Suspected Cancer

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for

Community-Based CPD Programming | Thompson 15 September, 2023

#### Case1: Tania, age 35

- Tania presents on a humid Friday afternoon with two complaints: a lump in the right side of her neck and an itchy rash which she says is present everywhere but worst over her back. Aside from a bout of infectious mononucleosis which you diagnosed five years ago, Tania had been in good health prior to the onset of these symptoms. She reports having lost about 7 lbs. in the past three weeks. Tania thinks her diffuse itching is from the stress of her job as inspector at a meat packaging plant. She thinks the hot weather over the past few weeks is also bothering her. She tends to sweat a lot at night, and feels restless when she tries to sleep.
- A rounded swelling in the vicinity of the anterior border of the mid-portion of her right sternomastoid is evident. Ovoid, firm, smooth and non-tender, it measures about 3 cm in its vertical axis. The horizontal diameter is difficult to assess because the lateral portion of the node seems to disappear under the muscle belly of the sternomastoid.

#### Lymphadenopathy

- Lymphadenopathy (lymph nodes that are abnormally large or have an abnormal consistency) estimated annual incidence of 0.6%.<sup>1</sup>
- Malignancy is the cause of unexplained lymphadenopathy in approximately 1.1% of cases in the general population, increasing significantly with age.<sup>2</sup>

<sup>1.</sup> Gaddey HL, Riegel AM. Unexplained lymphadenopathy: evaluation and differential diagnosis. Am Fam Physician 2016;94(11):896–903.

<sup>2.</sup> Fijten GH, Blijham GH. Unexplained lymphadenopathy in family practice. An evaluation of the probability of malignant causes and the effectiveness of physicians' workup. J Fam Pract 1988;27(4):373–6

## QUESTION #1: What questions do you have for Tania?

- Tania denies having recently had a sore throat, fever, coryza, or cough.
- She is unaware of any sores in her mouth.
- She noticed the neck lump about two weeks ago; it has doubled in size since she noticed it. The node is not sore.
- Tania's energy has been mostly "OK", although she thinks that job-related stress has fatigued her just slightly in the past few weeks. She sleeps somewhat fitfully.
- The itching has been present almost continuously for the past three weeks.
- She has no known scabies contacts.
- Tania does not consume alcoholic beverages.
- Tania has no history of recent travel.
- Her parents are of Scottish and Ukrainian background; she grew up in the south end of Winnipeg; and she has no known history of exposure to tuberculosis.

# QUESTION #2: What should the physical examination include in a patient who presents with a lateral neck mass?

- A complete history and physical examination, including the following:
  - the patient's general appearance and vital signs;
  - A head and neck examination\*, with special attention to the upper aerodigestive tract. The examination of the head and neck of patients who present with a lateral neck mass should include inspection of the face and neck for skin lesions suggestive of skin cancers (melanoma, squamous or basal cell). Take care to examine the scalp and the nape of the neck, which are often overlooked. In addition to inspection of the lips, tongue, dentition, gums and pharynx, an examination of the oral cavity should include inspection and (if clinically warranted) gloved palpation of the mucosa of the buccal and lingual sulci; and the base of the tongue.
  - A cranial nerve examination;
  - A full body lymph node examination;
  - A full body skin examination with particular attention paid to findings suspicious for melanoma;
  - Lung auscultation;
  - Careful palpation of the abdomen, looking especially for masses and for hepato- and splenomegaly;

#### Case 1 continued:

 Tania's resting pulse is 96, her blood pressure (right arm sitting) 104/72, T 368, O<sub>2</sub> saturation 95% on room air. Her head & neck examination discloses the enlarged right neck node, which is nontender and slightly mobile, and a slightly ruddy complexion. The oropharynx appears normal. On careful examination you cannot appreciate any other enlarged nodes of the head, neck, axillae, epitrochlear areas or groins. Her rash consists only of scattered faint scratches over her trunk and limbs, and some larger and slightly inflamed excoriations of her upper back, sparing the areas beyond the reach of her fingers. Her cardiorespiratory and abdominal examinations disclose no other abnormalities.

# QUESTION #3: What is on the differential diagnosis of a lateral neck mass?

- LATERAL NECK MASSES FALL BROADLY INTO THREE CATEGORIES: INFLAMMATORY; CONGENITAL; AND NEOPLASTIC. THESE COULD INCLUDE:
- Benign reactive adenopathy, e.g. from a viral or bacterial upper respiratory infection
- Lymphomas, both Hodgkin and NHL
- Nodal involvement from a primary cancer of the upper aerodigestive tract
- Nodal involvement from a primary cancer of thyroid or salivary glands
- Other primary tumors of the head and neck (e.g. Schwannoma, paraganglioma)
- Tuberculous lymphadenopathy of the neck ("scrofula")
- Other bacterial lymphadenitis (e.g. Staph aureus, Streptococcal, tularemia)
- HIV
- nodal spread from a skin cancer of the head or neck
- metastatic spread from a lung cancer
- branchial cleft cysts
- hemangiomas and vascular malformations

#### Case 1 continued:

 you are especially concerned, in view of Tania's cervical lymphadenopathy and constitutional symptoms, and diffuse pruritus, that she might have a malignancy. You are eager to obtain a tissue diagnosis.

# QUESTION #4: How should you go about obtaining A TISSUE DIAGNOSIS?

• In the workup of an as-yet undiagnosed pathological lymph node of the lateral neck, it is recommended that a fine-needle aspirate (FNA) be obtained prior to proceeding (in particular) with an excisional biopsy. This is done to rule out a primary squamous carcinoma of the head and neck. Excision of a nodal neck metastasis from a squamous carcinoma of the head or neck, if performed outside the context of definitive surgical management, can make subsequent surgery more extensive and difficult and can also increase the risk for recurrence of the disease in the contralateral neck. FNA biopsy of enlarged neck nodes is usually done by the consulting ENT surgeon or Head and Neck surgical oncologist; primary care clinicians, if comfortable with the procedure, can also perform FNA biopsy of suspicious neck nodes, but should not delay referral to a consultant (for example, pending return of the pathology report, or for the sake of scheduling a second visit for the procedure).

#### Case 1 continued:

- You proceed with an FNA of the enlarged neck node on the occasion of that visit. Anticipating the possible need for an excisional biopsy of the node to confirm your suspicion of a malignancy, and concerned about her 'B' symptoms, you dictate a referral letter to a Head & Neck Surgeon, explaining the urgency of the matter to your clinic administrative staff, who promise to get the referral 'out the door in 24'. To further expedite the referral, you telephone the consultant, who advises you he will see your patient within the next 14 days\*.
- \*visit CancerCare Manitoba's website at <a href="http://www.cancercare.mb.ca/diagnosis">http://www.cancercare.mb.ca/diagnosis</a> and click on the various disease- specific *In Sixty* Pathways to see the recommended timelines pertinent to the work-up of a suspected cancer.

# QUESTION #5: WHAT OTHER WORK UP SHOULD YOU UNDERTAKE WHILE WAITING FOR THE EXCISIONAL BIOPSY?

- You should provide Tania with requisitions for a chest X-ray, a CBC and HIV serology to be done that same day. If the lymphocyte count is elevated on the CBC, you should also order flow cytometry.
- Renal function tests and LFTs should be obtained
- Send an urgent requisition for computed tomography of the neck, chest, abdomen & pelvis. Ensure that the report is copied to the Head & Neck surgeon and to any other consultant to whom you refer your patient.
- Offer your patient a referral to Cancer Navigation services at your local/regional Hub. A Nurse Navigator can answer your patient's questions about Manitoba's cancer care system, track your patient's progress through the system, and arrange for tissue pathology reports to be forwarded to you and to the specialists involved in your patient's care.

#### Case 1 continued:

- The chest radiograph report arrives early the next week. It suggests possible mediastinal lymphadenopathy. You telephone the radiologist at the community hospital to which the CT requisition was sent; he agrees to expedite the appointment for the CT scans you have requested.
- Tania comes back to see you for review the next week, two weeks after her initial visit with you. The FNA showed numerous lymphocytes but was not adequate for diagnosis. She has already had an excisional biopsy of the enlarged neck node. You telephone the Head & Neck Surgeon, who says he has just received the pathology report which confirms the suspected diagnosis of Hodgkin lymphoma. The CT report, just back this morning, mentions two other enlarged nodes in the neck, including a posterior cervical node and a supraclavicular node, both on the right. Bulky mediastinal lymphadenopathy is also remarked, including a node which impinges on the superior vena cava. Tania notes that along with the work stress she has felt restless at night, she is having insomnia, and she feels particularly agitated if she lies down, so she has been sleeping in her recliner for the past week. As Tania takes a seat in your examination room, you are struck by the ruddiness of her complexion.

# QUESTION #6: WHAT CONDITION IS COMPLICATING TANIA'S NEWLY DIAGNOSED HODGKIN LYMPHOMA? HOW WILL YOU MANAGE IT?

- Tania has an evolving superior vena cava syndrome. Depending on Tania's clinical status and your practice setting, you might send her directly to the nearest ER, or arrange through the on-call hematologist for Tania's admission that same day to a tertiary care facility where she can be given high dose steroids, with a view to starting chemotherapy in the next 24-48 hours.
- Radiotherapy is also a treatment that can be given for SVC syndrome if urgent initiation of chemotherapy is not feasible.
- Send to CCMB along with the written (urgent) hematology-oncology request all relevant history, lab data, and imaging and pathology reports.
- There are no proven connections between environmental exposures and Hodgkin Lymphoma, but an increase in incidence has been reported in woodworkers, farmers, and meat workers.
- About 70% of Hodgkin lymphoma cases present with cervical adenopathy alone, with no associated B-symptoms.

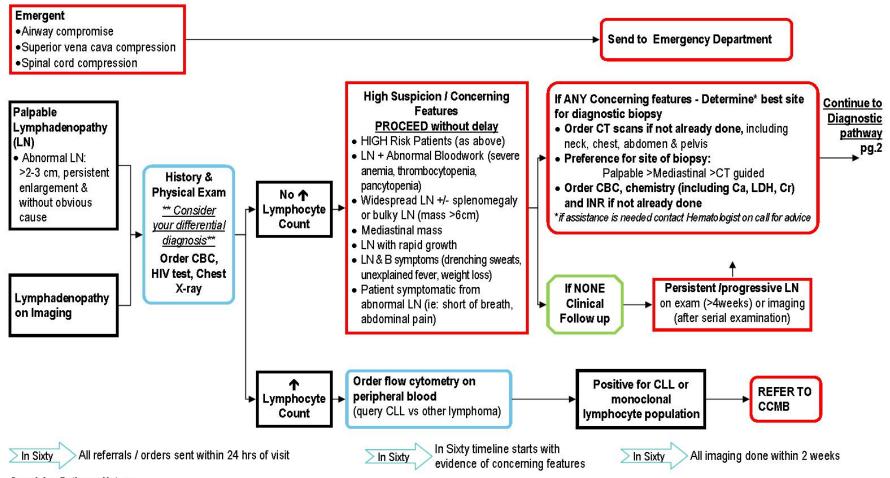


#### Work-Up of Lymphadenopathy Suspicious for LYMPHOMA

Timeline and Legend pg.5

RISK FACTORS: HIGH risk: immune deficiency (ie. HIV or organ transplant), autoimmune disease +/- immune suppressing medications, and history of lymphoma PRACTICE POINTS: \*\*Consider your differential diagnosis\*\* including reactive LN due to infection/inflammation, metastatic malignancy, and autoimmune disease. This document applies to adults 17 years of age or older.

PRACTICE POINTS: All referrals sent within 24 hrs of visit. Provide complete information as requested to avoid delays. Ensure patient and family is well informed and receives appointment information. If patient is in distress, offer referral to local counsellor. See Supporting Information for Clinicians (pg 4) for contacts and resources. Contact the Cancer Question Helpline for Primary Care for assistance.



#### Suspicion Pathway Notes:

#### Case 2: Helen Bighetty

- Helen Bighetty is a 58 year old female, well known to you, presenting today with a recurrence of low back pain she has had on & off for 10+ years.
- Helen has hypertension diagnosed with a home blood pressure mean BP of 138/86
- treated with amlodipine 5 mg daily.
- 35 pack-year history of cigarette smoking.
- slightly overweight.
- Cholecystectomy age 36.
- NKA

- "I've thrown my back out again."
- What do you want to know?
- Her pain in mid-to-low back (higher in her back than usual)
- Past 5 weeks
- Wakes with pain
- Partial relief with acetaminophen 650 mg QID +/- ibuprofen up to 400 mg TID
- No leg pain or paraesthesias
- Denies fecal incontinence or urinary retention

#### On examination:

- LS spine ROM within normal for age
- No provocation of pain with flexion or extension
- Punch tenderness at ~ T12 and L2
- SLR negative bilaterally
- Lower limb power and sensation normal
- Anal sphincter tone normal
- Perianal sensation normal.

#### What imaging do you want?

- Plain radiographs of T, LS spine
- Lytic lesions at T12 and L1,2 & 3 vertebral bodies



https://www.stritch.luc.edu/lumen/meded/radio/curriculum/neurology/spine\_plain\_xray\_2013.htm

#### Lytic Spine Lesions

- FOG MACHINES
- F: fibrous dysplasia (FD) or fibrous cortical defect (FCD)
- O: osteoblastoma
- **G**: giant cell tumor (GCT) or geode
- M: metastasis(es)/myeloma
- A: aneurysmal bone cyst (ABC)
- C: chondroblastoma or chondromyxoid fibroma
- **H:** hyperparathyroidism (brown tumor)
- I: infection (osteomyelitis) or infarction
- **N**: non-ossifying fibroma (NOF)
- E: enchondroma or eosinophilic granuloma (EG)
- **S:** simple (unicameral) bone cyst

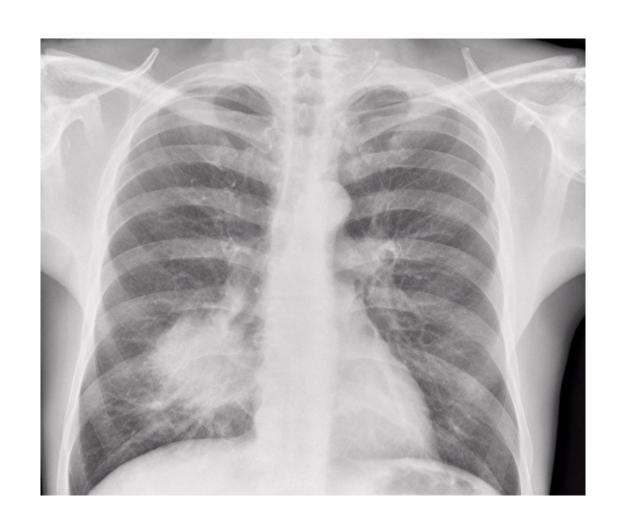
Hakim T, Ashraf A, Sharma R, et al. Lucent/lytic bone lesion - differential diagnosis (mnemonic). Reference article, Radiopaedia.org (Accessed on 24 Aug 2023) https://doi.org/10.53347/rID-450

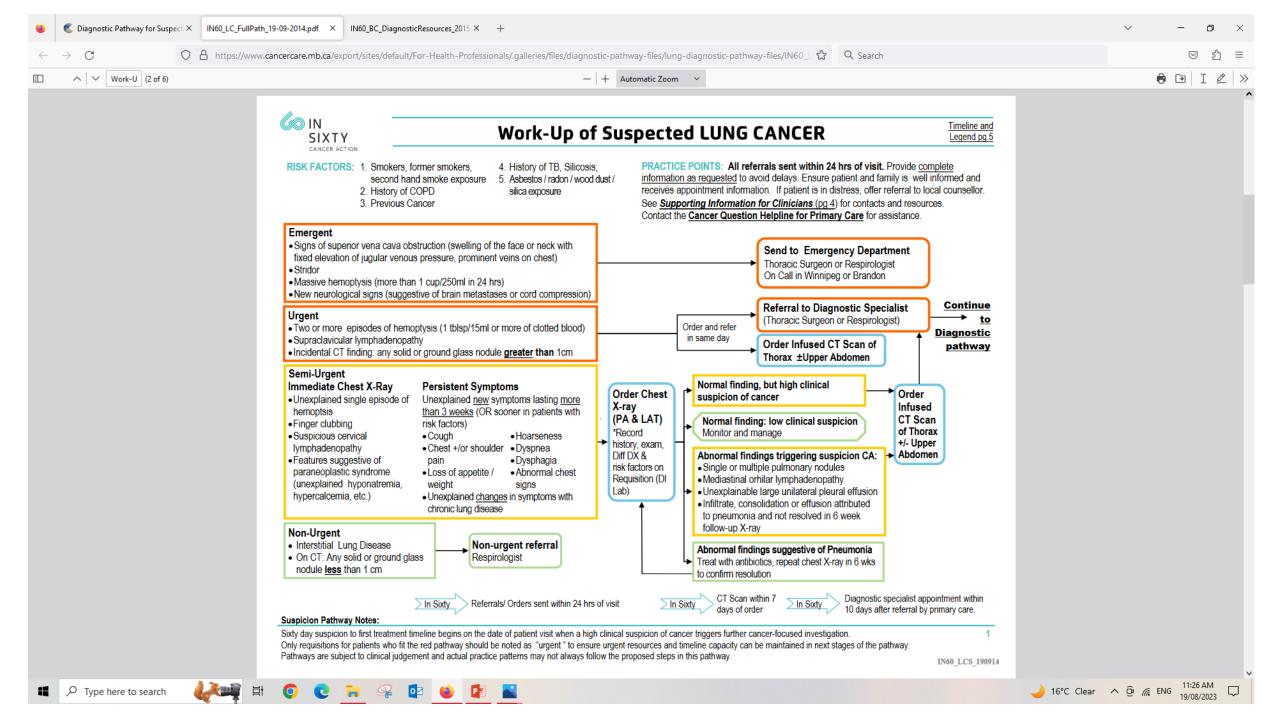
#### What further imaging do you want?

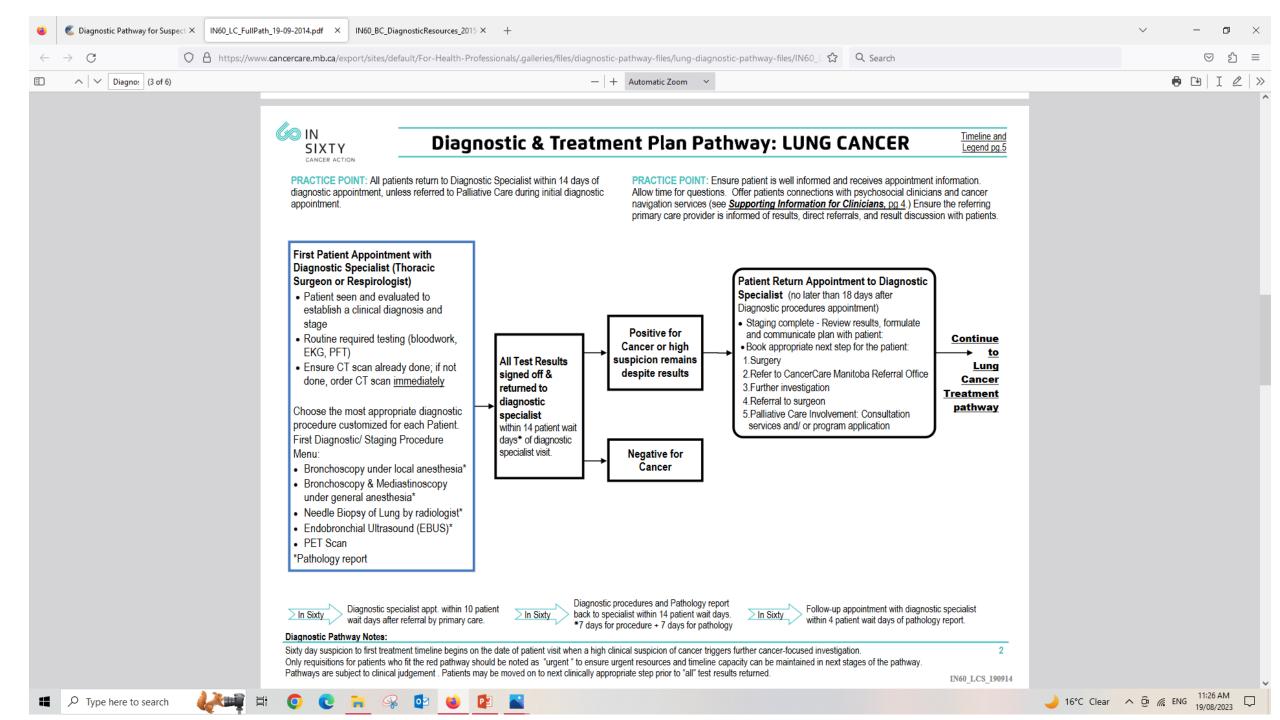
- CT: T, LS spine
- Chest radiograph

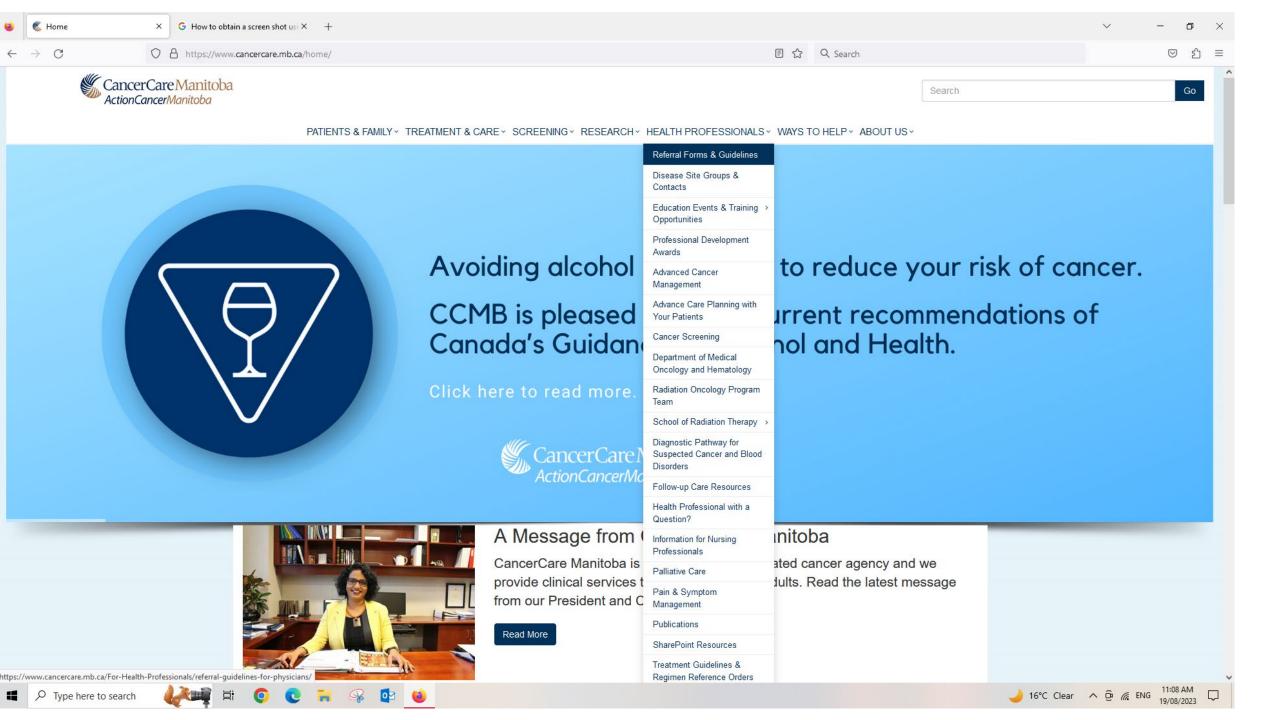
#### What lab work do you want?

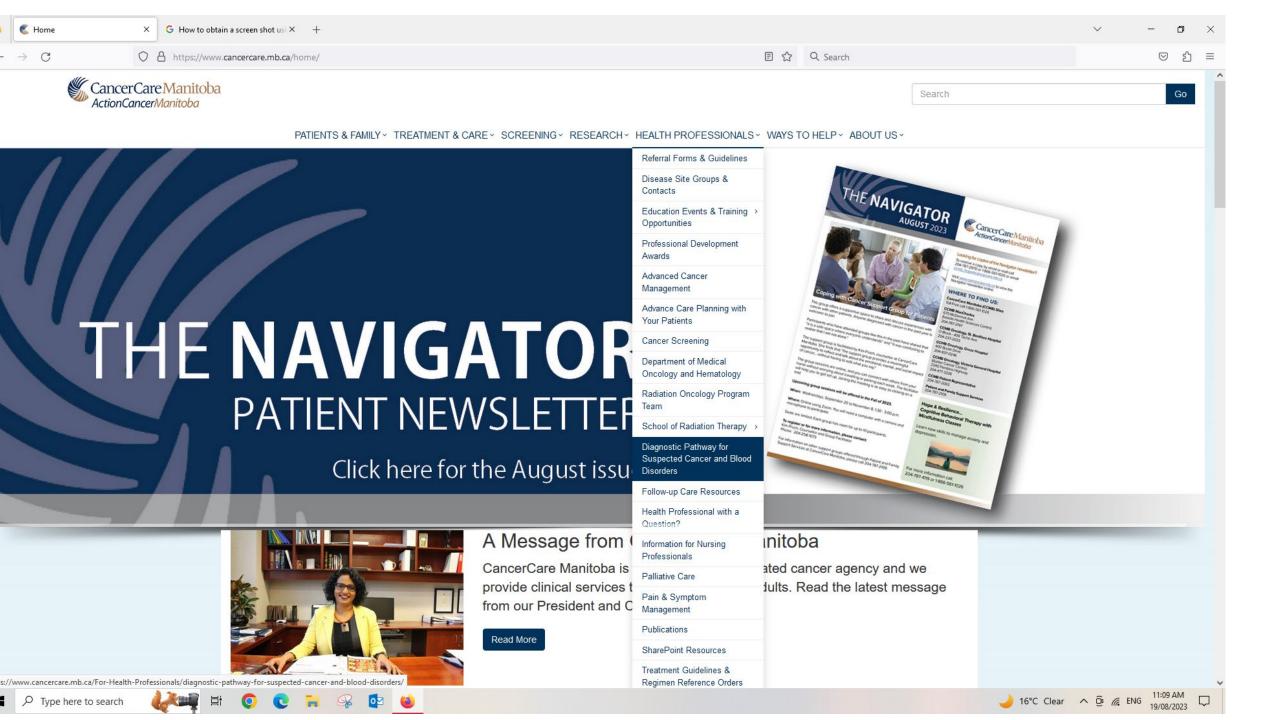
- CBC, SPEP, FLRCR, serum Cr, extended 'lytes, liver enzymes.
- U/A for blood

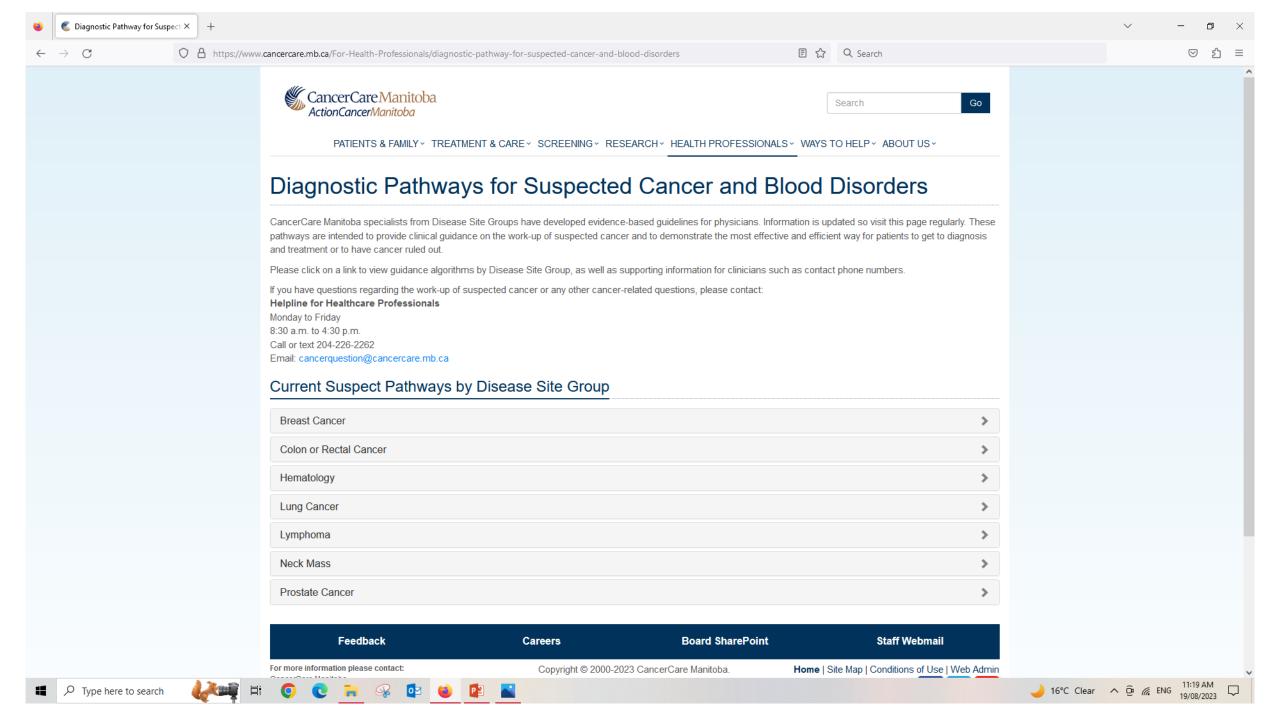


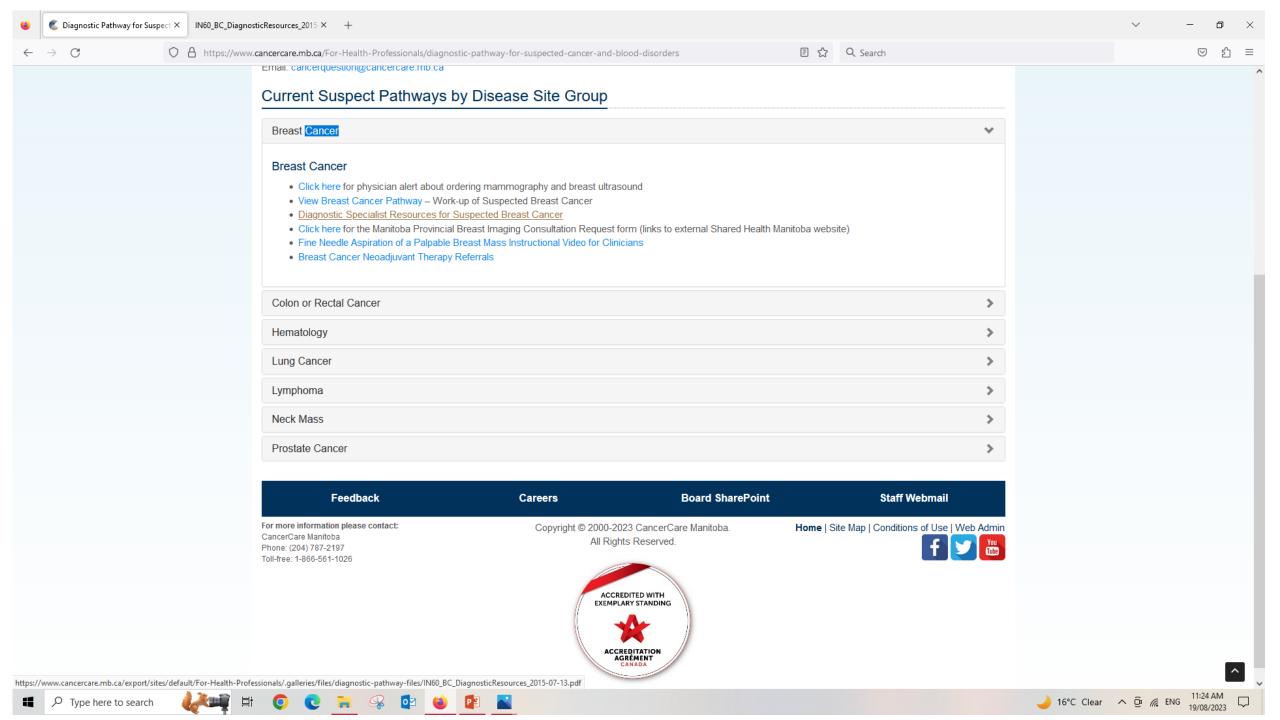




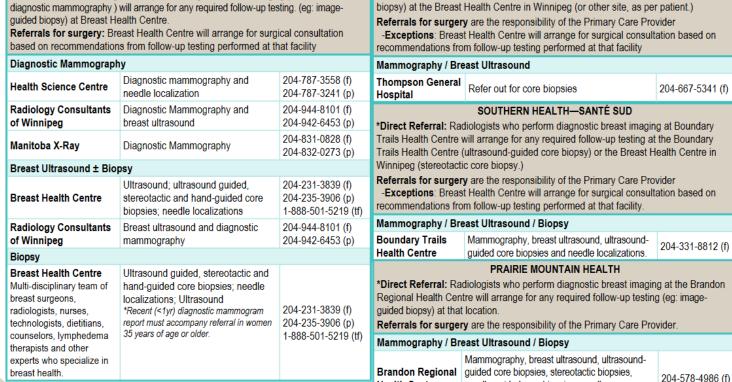












**Health Centre** 

needle-guided core biopsies, needle localizations, breast MRI & MR-guided biopsies.



From Suspicion of Cancer to Treatment In Sixty Days

For Review: IN60 BCS 2015-07-13



















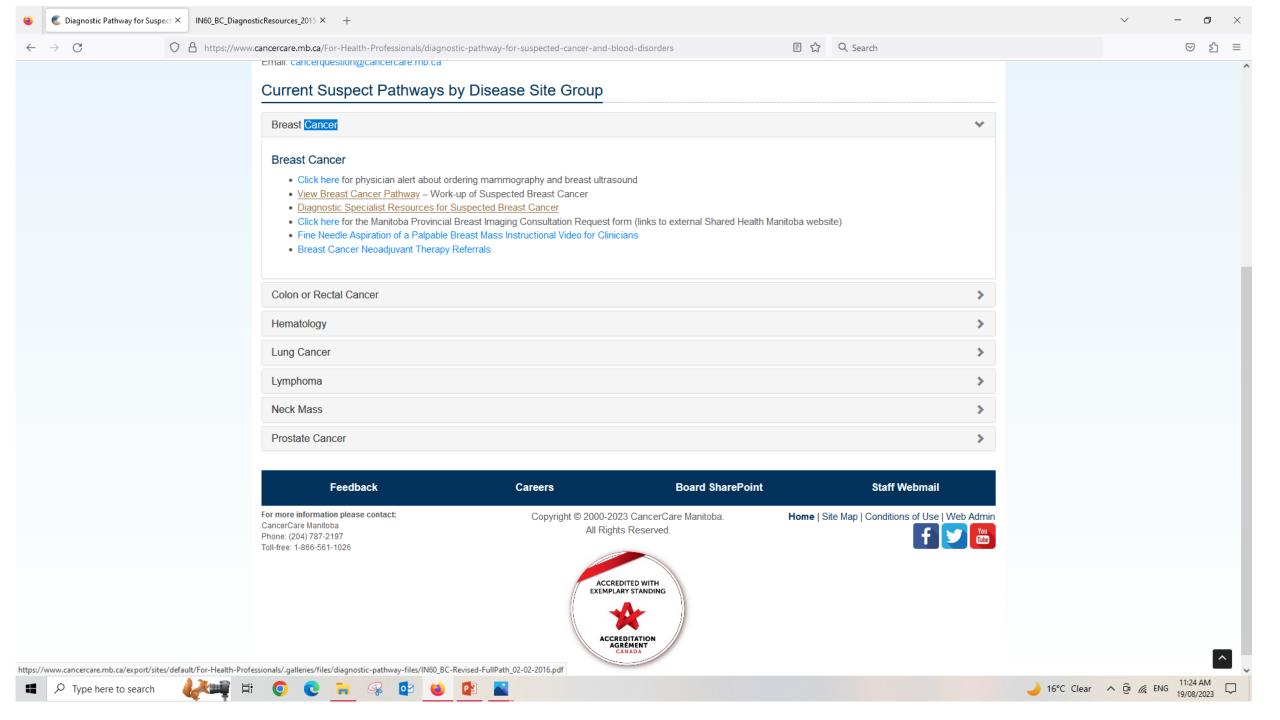


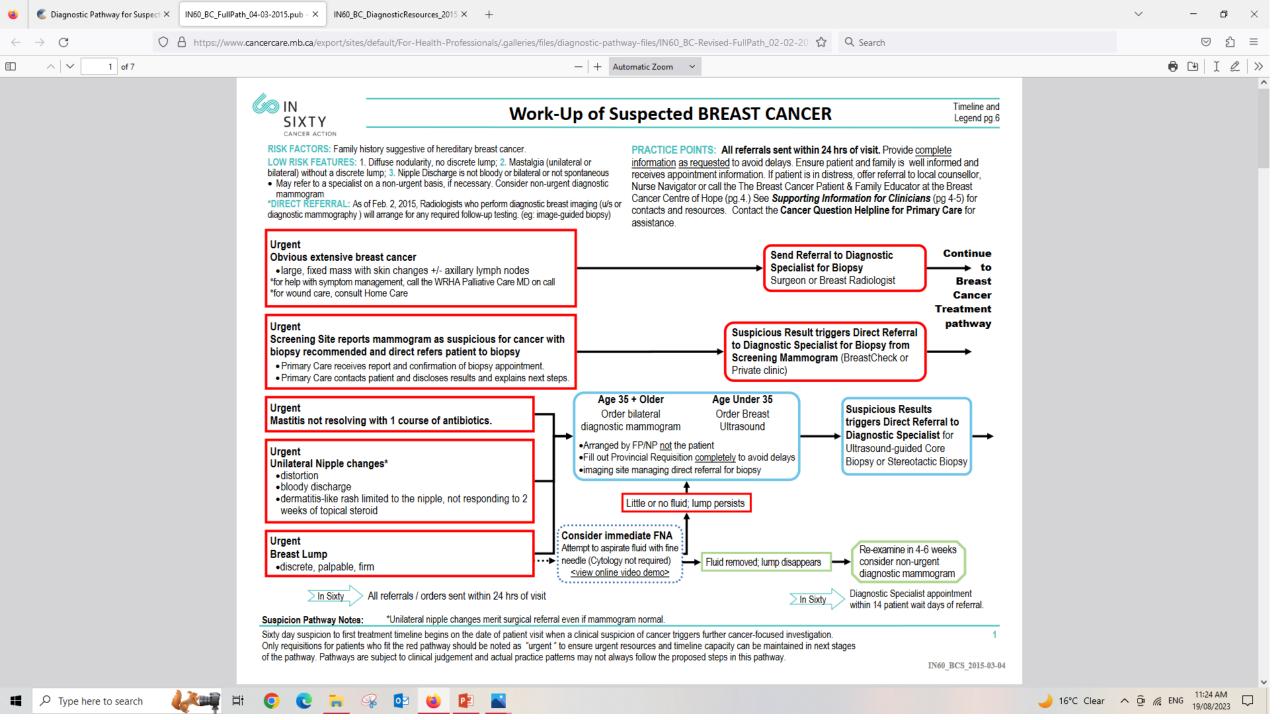


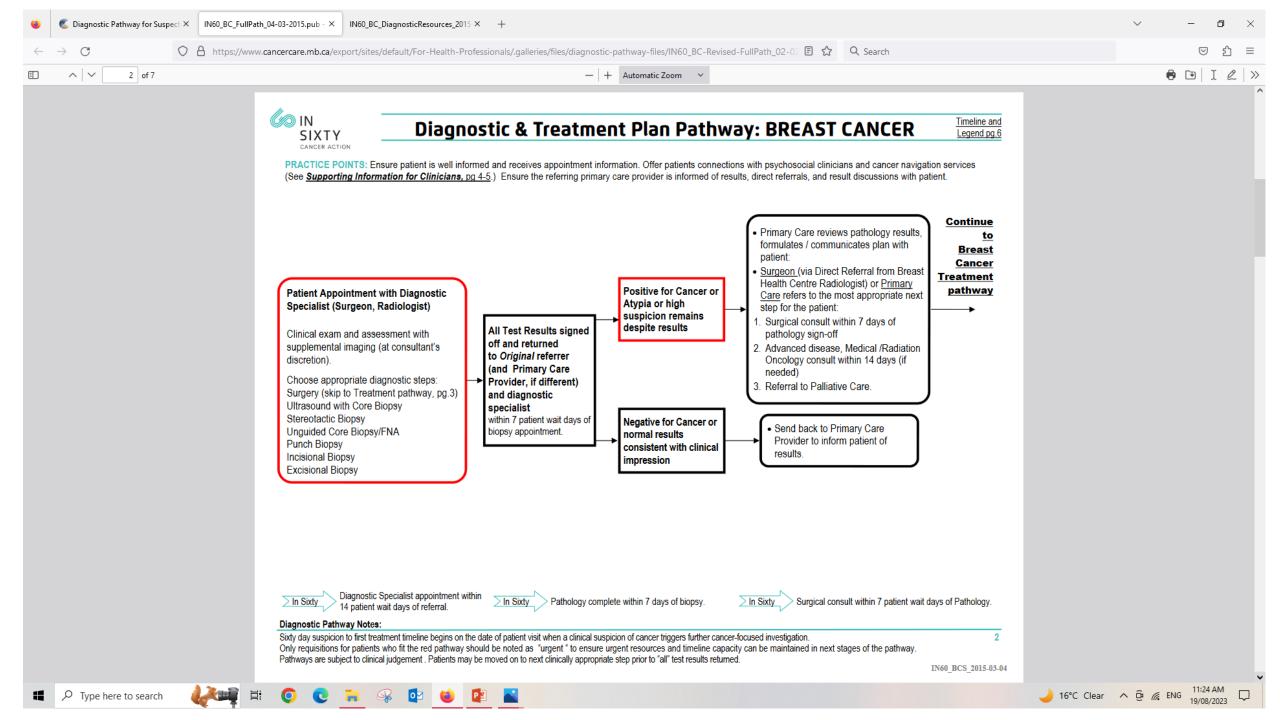












#### Suspicious?

- DVT/PE
- Lymphadenopathy
- Cutaneous signs of malignancy
- B-symptoms

#### Pretest probability of deep vein thrombosis (Wells score)

Clinical feature	Score
Active cancer (treatment ongoing or within the previous six months or palliative)	1
Paralysis, paresis, or recent plaster immobilization of the lower extremities	1
Recently bedridden for more than three days or major surgery, within four weeks	1
Localized tenderness along the distribution of the deep venous system	1
Entire leg swollen	1
Calf swelling by more than 3 cm when compared to the asymptomatic leg (measured below tibial tuberosity)	1
Pitting edema (greater in the symptomatic leg)	1
Collateral superficial veins (nonvaricose)	1
Alternative diagnosis as likely or more likely than that of deep venous thrombosis	-2
Score	
High probability	3 or greater
Moderate probability	1 or 2
Low probability	0 or less
Modification:	
This clinical model has been modified to take one other clinical feature into account: a previously documented deep vein thrombosis (DVT) is given the score of 1. Using this modified scoring system, DVT is either likely or unlikely, as follows:	
DVT likely	2 or greater
DVT unlikely	1 or less

#### Adapted from:

- Wells PS, Anderson DR, Bormanis J, et al. Value of assessment of pretest probability of deep-vein thrombosis in clinical management. Lancet 1997; 350:1795
- Wells PS, Anderson, DR, Rodger M, et al. Evaluation of D-dimer in the diagnosis of suspected deep-vein thrombosis. N Engl J Med 2003; 349:1227.

### DVT/PE – work up for malignancy?

- No benefit from working a patient up for an occult cancer in the setting of a provoked DVT
- A limited work up for occult malignancy is justified in patients presenting with an unprovoked first DVT/PE

## DVT/PE – work up for malignancy?

**Limited** — The components of a limited testing strategy in patients with a first episode of uncomplicated unprovoked VTE include the following:

- •A complete history for the signs and symptoms of malignancy
- •A thorough physical examination, including digital rectal examination, testing for fecal occult blood, and pelvic examination in women
- •Basic laboratory testing including complete blood count (CBC) and smear, ESR, urinalysis, electrolytes, calcium, creatinine, and liver function tests
- Routine age-appropriate cancer screening
- Chest radiograph

#### DVT/PE – work up for malignancy?

- In patients who present with an unprovoked first DVT/PE and are at higher risk for having an occult malignancy, a more extensive work up is suggested
- Investigations are frequently symptom-directed. They may include but are not limited to the following:
- All the components of a limited strategy
- A chest, abdominal, and pelvic CT scan
- Tumor markers (carcinoembryonic antigen, alpha-fetoprotein, CA 19-9, CA 125, PSA)
- Mammography and a Papanicolaou smear in women
- Upper and lower gastrointestinal tract evaluation (eg, endoscopy or CT colonography)

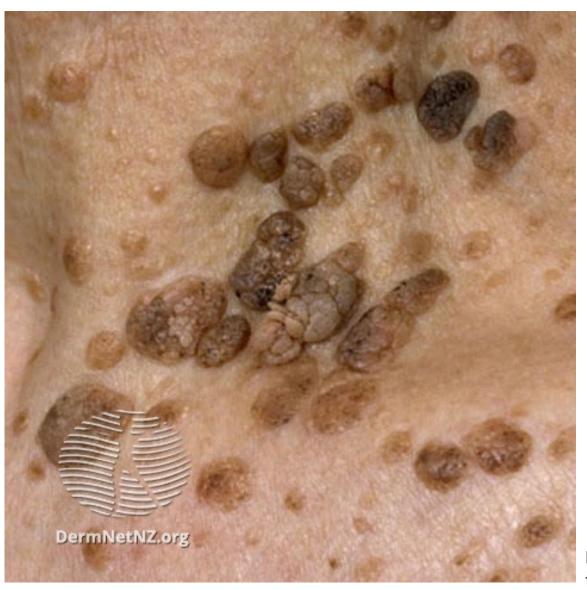
UpToDate: Evaluating adult patients with established venous thromboembolism for acquired and inherited risk factors

#### Cutaneous Disorders Associated with Cancer

#### Sign of Leser-Trélat

- Multiple seborrheic keratoses
- often pruritic
- associated with internal malignancies, esp. GI adenocarcinomas:
- Gastric
- Liver
- Colorectal
- Pancreatic

## Sign of Leser- Trelat



https://dermnetnz.org/topics/sign-of-leser-trelat

#### Cutaneous Disorders Associated with Cancer

#### **Acanthosis Nigricans**

- Hyperkeratotic
- Velvety
- Darkly pigmented plaques
- Benign etiology most common (insulin resistance, obesity)
- Younger age of onset
- Confined to flexural creases (neck, axillae, groins)
- Malignant AN more rapidly progressive
- More extensive mucosal membranes, palms & soles
- elderly
- Gastric Ca (+ other intraabdominal)

## Acanthosis Nigricans



https://dermnetnz.org /images/acanthosisnigricans-images

## Cutaneous Disorders Associated with Cancer <u>Sweet Syndrome</u>



## Cutaneous Disorders Associated with Cancer <u>Sweet Syndrome</u>





## Cutaneous Disorders Associated with Cancer <u>Sweet Syndrome</u>

