流感臨床診斷 妈 一

成大醫院感染科 羅景霳 20250726



- 流感臨床診斷與檢驗

臨床表現

- 流感臨床診斷與檢驗

臨床表現

診斷工具

流感臨床診斷與檢驗

臨床表現

診斷工具

鑑別診斷

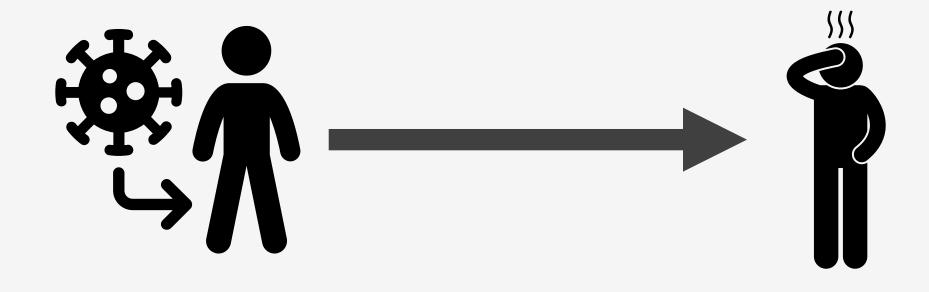
流感臨床診斷與檢驗

臨床表現

診斷工具

鑑別診斷

- 流感的潛伏期



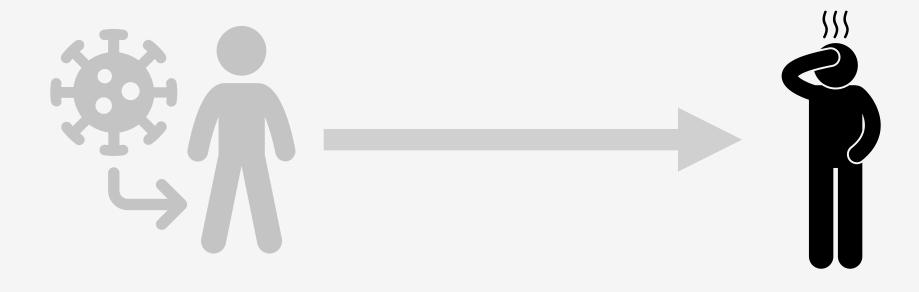
- 流感的潛伏期



群聚個案的時間間隔

大約是3~4天

- 流感的潛伏期



- 流感的臨床表現

- 流感的臨床表現

Uncomplicated

- 流感的臨床表現

- Uncomplicated
- Complicated

- 流感的臨床表現
 - Uncomplicated
 - Complicated

- Typical symptom of uncomplicated influenza
 - Respiratory symptom
 - Systemic symptom

- Typical symptom of uncomplicated influenza
 - Respiratory symptom
 - Systemic symptom
 - Non-respiratory symptom

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Respiratory symptom

Dry cough

- Dry cough
- Severe pharyngeal pain

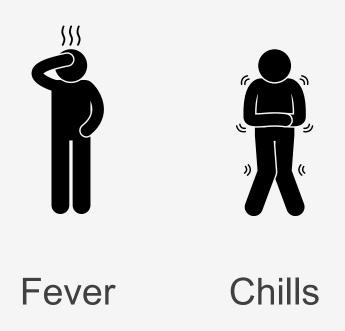
- Dry cough
- Severe pharyngeal pain
- Nasal obstruction and discomfort

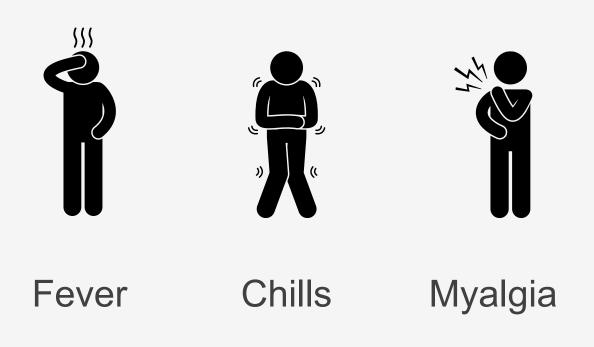
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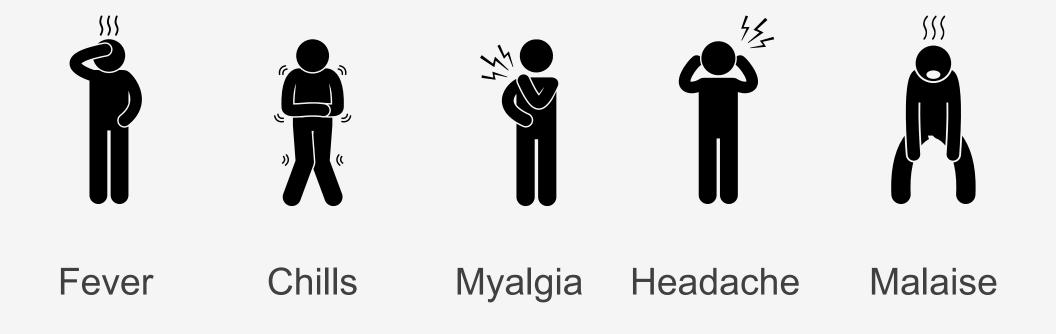
Systemic symptom



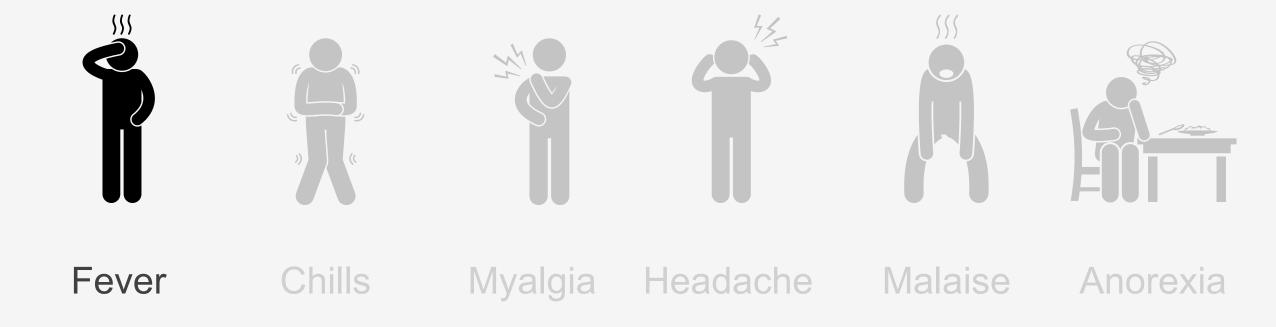












Systemic symptom

Abrupt onset



Systemic symptom

- Abrupt onset
 - 知道明確時間



Systemic symptom

- Abrupt onset
 - 知道明確時間



• $37.8 \sim 40.0^{\circ}C (41.1^{\circ}C)$

Systemic symptom

- Abrupt onset
 - 知道明確時間



- $37.8 \sim 40.0^{\circ}C (41.1^{\circ}C)$
- Highest on D1

Systemic symptom

- Abrupt onset
 - 知道明確時間



- $37.8 \sim 40.0^{\circ}C (41.1^{\circ}C)$
- Highest on D1
 - 0.3~0.6°C lower afterward

Systemic symptom

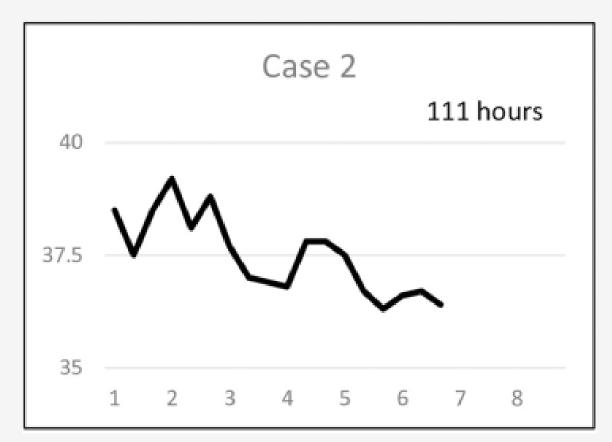
- Abrupt onset
 - 知道明確時間



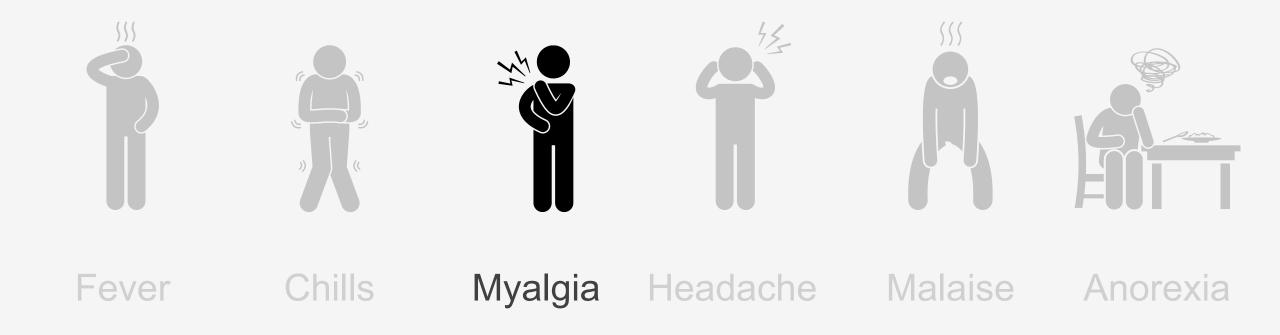
- $37.8 \sim 40.0^{\circ}C (41.1^{\circ}C)$
- Highest on D1
 - 0.3~0.6°C lower afterward
 - Duration: 3 days (4~8 days)

- Fever curve of influenza





Typical symptom of uncomplicated influenza Systemic symptom



Systemic symptom

Most troublesome



Systemic symptom

- Most troublesome
- Relate to height of fever



Systemic symptom

Range



Systemic symptom

- Range
 - Extremities



Systemic symptom

- Range
 - Extremities
 - Long muscle of back



Systemic symptom

- Range
 - Extremities
 - Long muscle of back
 - Eye muscle



Systemic symptom

- Range
 - Extremities
 - Long muscle of back
 - Eye muscle
 - pain while gazing laterally



Systemic symptom

- Range
 - Extremities
 - Long muscle of back
 - Eye muscle
 - pain while gazing laterally





- Typical symptom of uncomplicated influenza
 - Respiratory symptom
 - Systemic symptom
 - Non-respiratory symptom

Non-respiratory symptom

Gastrointestinal symptom

Non-respiratory symptom

- Gastrointestinal symptom
 - Vomiting
 - Diarrhea

Non-respiratory symptom

- Gastrointestinal symptom
 - Vomiting
 - Diarrhea
- More in children (10~20%)



Milder systemic symptom

- Atypical symptom of uncomplicated influenza
 - Milder systemic symptom
 - No fever
 - No myalgia

- Atypical symptom of uncomplicated influenza
 - Milder systemic symptom
 - No fever
 - No myalgia
 - Only generalized symptom

- Atypical symptom of uncomplicated influenza
 - Milder systemic symptom
 - No fever
 - No myalgia
 - Only generalized symptom
 - Anorexia/Malaise/Weakness
 - Dizziness

- Milder systemic symptom
 - No fever
 - No myalgia
- Only generalized symptom
 - Anorexia/Malaise/Weakness
 - Dizziness
 - Altered mental status (Older adults)

Who would have Atypical symptom of uncomplicated influenza

Who would have Atypical symptom of uncomplicated influenza

Age > 65 years old

Who would have Atypical symptom of uncomplicated influenza

- Age > 65 years old
- Immunocompromised

Some respiratory symptom would persist after systemic symptom diminish

Some respiratory symptom would persist after systemic symptom diminish

- Recurrent cough
- Hoarseness
- Dry/sore throat

Some respiratory symptom would persist after systemic symptom diminish

- Recurrent cough
- Hoarseness
- Dry/sore throat

不是變嚴重

- 流感的臨床表現
 - Uncomplicated
 - Complicated



≥ 65 y/o





≥ 65 y/o

< 5 y/o

< 2 y/o







< 5 y/o

< 2 y/o



Pregnant /2wk postpartum







< 5 y/o

< 2 y/o



Pregnant /2wk postpartum



Nursing home Long term care







< 5 y/o

< 2 y/o



Pregnant /2wk postpartum



Nursing home Long term care



Medical condition







< 5 y/o





Pregnant /2wk postpartum



Nursing home Long term care



Medical condition

Medical condition associated with complication of influenza

- Neurological and neurodevelopmental conditions
- Asthma
- Chronic lung disease
- Heart disease
- Kidney disorder
- Liver disorder

- Endocrine disorders
- Metabolic disorder
- BMI ≥ 40
- Blood disorder
- Weakened immune system
- Children<19 y/o under long-term ASA

-Complication of influenza

- Complication of influenza

Respiratory

- Complication of influenza

- Respiratory
- Extrapulmonary

-Complication of influenza

- Respiratory
- Extrapulmonary

-Respiratory complication of influenza

Pneumonia

Symptom of pneumonia

- Cough with dyspnea
- Tachypnea
- Hypoxia
- Fever

Primary influenza viral pneumonia

- Primary influenza viral pneumonia
- Secondary bacterial pneumonia

- Primary influenza viral pneumonia
- Secondary bacterial pneumonia
- Mixed viral and bacterial pneumonia

- Primary influenza viral pneumonia
 - Persistent fever and symptom after 3-5 days of symptom

- Secondary bacterial pneumonia
 - Improvement of influenza symptom
 - Relapse of fever and cough with purulent sputum

- Mixed viral and bacterial pneumonia
 - Gradual progression
 - or Transient improvement followed by worsening

-Complication of influenza

- Respiratory
- Extrapulmonary

- Extrapulmonary complication of influenza -
 - Cardiac
 - Central nervous system
 - Myositis and rhabdomyolysis
 - Multisystem organ failure
 - Concomitant infection

- Extrapulmonary complication of influenza -
 - Cardiac
 - Central nervous system
 - Myositis and rhabdomyolysis
 - Multisystem organ failure
 - Concomitant infection

Extrapulmonary complication of influenza

Cardiac

- Myocardial infarction
- Myocarditis/Pericarditis

- Extrapulmonary complication of influenza
 - Central nervous system
 - Guillain-Barre syndrome

- Guillain-Barre syndrome

- Incidence of GBS after influenza
 - 30 days: 16.64 (95% CI: 9.37-29.54)
 - 90 days: 7.35 (95% CI: 4.36-12.38)

- Guillain-Barre syndrome

- Incidence of GBS after influenza
 - 30 days: 16.64 (95% CI: 9.37-29.54)
 - 90 days: 7.35 (95% CI: 4.36-12.38)
- Clinical presentation
 - Progressive and symmetric muscle weakness
 - Absent or depressed deep tendon reflexes

- 臨床表現・

- 在流感流行期, 臨床診斷的正確率高
- Uncomplicated influenza中,全身症狀, 如發燒和肌肉痠痛等,是流感的特色
- Influenza complication中,肺炎是最常見的併發症,如果病程超出預期,要特別注意

流感臨床診斷與檢驗

臨床表現

診斷工具

鑑別診斷

-Case



42 years old

Case



42 years old

Cough with sputum and rhinorrhea for one week Fever with chillness for 4 days

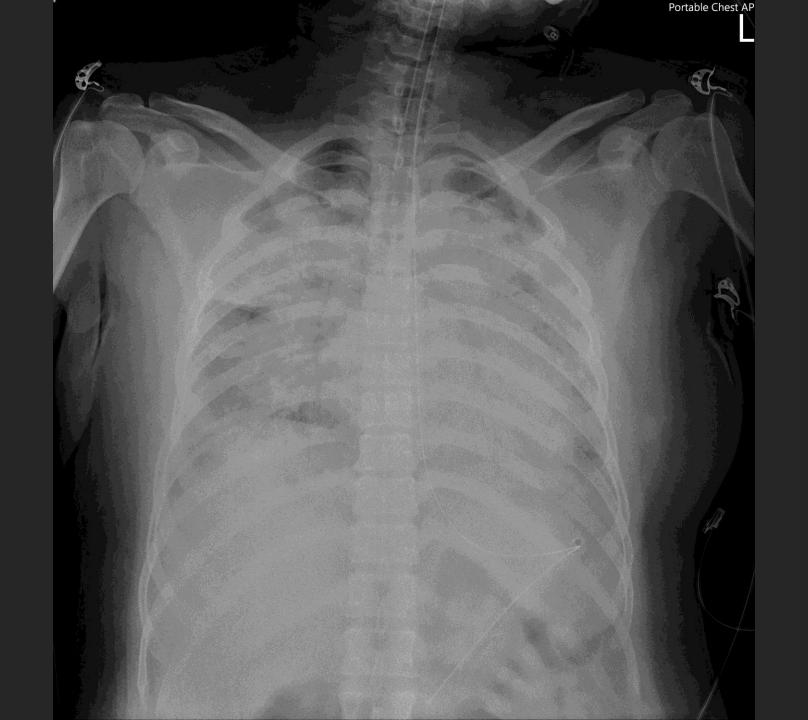
Case



42 years old

Cough with sputum and rhinorrhea for one week Fever with chillness for 4 days

- Diarrhea



-Case



42 years old

法定傳染病通報送檢檢驗報告

通報疾病: SARS-CoV-2病毒檢測 檢體種類: 鼻咽拭子/咽喉拭子

病原體檢驗方法:螢光定量聚合酶連鎖反應(real-time PCR)

檢驗結果: 陰性

綜合檢驗結果: 陰性

檢驗單位:國立成功大學醫學院附設醫院

-Case



42 years old

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Influenza virus type A+B antigen(酵素免疫法)(快診-流行感冒病毒A+B型抗原檢驗(酵素免疫法))
Influenza virus type A antigen Negative
Influenza virus type B antigen Negative

排除流感?

Why to test

- Why to test
- Whom to test

- Why to test
- Whom to test
- What to test

- Why to test
- Whom to test
- What to test
- How to test

- Why to test
- Whom to test
- What to test
- How to test

-Why to test

 Reduce unnecessary testing for other etiologies

Why to test

- Reduce unnecessary testing for other etiologies
- Reduce inappropriate antibiotic use

-Why to test

- Reduce unnecessary testing for other etiologies
- Reduce inappropriate antibiotic use
- Improve the effectiveness of infection prevention and control measures

-Why to test

- Reduce unnecessary testing for other etiologies
- Reduce inappropriate antibiotic use
- Improve the effectiveness of infection prevention and control measures
- Increase the appropriate use of antiviral medications

不能因為要等檢驗

而延遲抗病毒藥使用

- 流感診斷工具

- Why to test
- Whom to test
- What to test
- How to test

If result will influence management

- If result will influence management
- Public health activity

- If result will influence management
- Public health activity

- If result will influence management
 - Antiviral or antimicrobial agents

- If result will influence management
 - Antiviral or antimicrobial agents
 - Further diagnostic evaluation

- If result will influence management
 - Antiviral or antimicrobial agents
 - Further diagnostic evaluation
 - Prophylaxis for high risk contacts

- If result will influence management
 - Antiviral or antimicrobial agents
 - Further diagnostic evaluation
 - Prophylaxis for high risk contacts
 - Infection control intervention

- If result will influence management
- Public health activity

- Public health activity
 - Interventions for outbreak management

- Influenza is circulating
- Influenza is not circulation

- Influenza is circulating
- Influenza is not circulation

Influenza is circulating

- Present influenza-like illness
 - Immunocompromised
 - high risk of complication

Influenza is circulating

- Present influenza-like illness
 - Immunocompromised
 - high risk of complication
- Acute respiratory symptoms
 - Exacerbation of chronic medical condition
 - Influenza complication

Influenza is circulating

- Present influenza-like illness
 - Immunocompromised
 - high risk of complication
- Acute respiratory symptoms
 - Exacerbation of chronic medical condition
 - Influenza complication
- Hospitalized patients
 - Acute respiratory symptoms
 - Exacerbations of chronic medical condition

流行期間

有風險/症狀惡化可檢驗

- Influenza is circulating
- Influenza is not circulation

Influenza is not circulating

Relevant epidemiologic exposure

Influenza is not circulating

- Relevant epidemiologic exposure
 - Exposure to person with influenza

Influenza is not circulating

- Relevant epidemiologic exposure
 - Exposure to person with influenza
 - Outbreak of respiratory illness of uncertain cause

Influenza is not circulating

- Relevant epidemiologic exposure
 - Exposure to person with influenza
 - Outbreak of respiratory illness of uncertain cause
 - Recent travel in an area with influenza activity

非流行期間

要有曝觸才要驗

- 流感診斷工具

- Why to test
- Whom to test
- What to test
- How to test

Molecular assay

- Molecular assay
- Antigen detection assay

- Molecular assay
- Antigen detection assay
- Others

- Molecular assay
- Antigen detection assay
- Others

Molecular assay is preferred if available

Molecular assay is preferred if available

- High sensitivity
- High specificity

Molecular assay is preferred if available

Molecular assay is preferred if available

Conventional RT-PCR

Molecular assay is preferred if available

- Conventional RT-PCR
- Rapid molecular tests

Molecular assay is preferred if available

- Conventional RT-PCR
- Rapid molecular tests
- Multiplex RT-PCR

Molecular assay is preferred if available

- Conventional RT-PCR
- Rapid molecular tests
- Multiplex RT-PCR

Conventional RT-PCR

Distinguish influenza A and B

Conventional RT-PCR

- Distinguish influenza A and B
- Influenza A subtype

Conventional RT-PCR

- Distinguish influenza A and B
- Influenza A subtype

Turnaround time: 1-8 hours

Rapid molecular tests

Distinguish influenza A and B

Rapid molecular tests

- Distinguish influenza A and B
- No influenza A subtype

Rapid molecular tests

- Distinguish influenza A and B
- No influenza A subtype

Turnaround time: 15~30 min

Multiplex RT-PCR

Several pathogens detection

-Filmarray respiratory panel





-Filmarray respiratory panel





- Adenovirus
- Human Rhinovirus/ Enterovirus
- Influenza virus
- Respiratory syncytial virus
- Parainfluenza
- Human metapneumovirus
- Coronavirus
 - SARS-CoV-2

-Filmarray respiratory panel



- Adenovirus
- Human Rhinovirus/ Enterovirus
- Influenza virus
- Respiratory syncytial virus
- Parainfluenza
- Human metapneumovirus
- Coronavirus
 - SARS-CoV-2



- Chlamydia pneumoniae
- Bordetella pertussis
- Bordetella parapertussis
- Mycoplasma pneumoniae

Multiplex RT-PCR

• 貴又容易被核刪

- 貴又容易被核刪
- 哪些族群建議檢測

- 貴又容易被核刪
- 哪些族群建議檢測
 - Immunocompromised

- 貴又容易被核刪
- 哪些族群建議檢測
 - Immunocompromised
 - Requiring hospitalization

- 貴又容易被核刪
- 哪些族群建議檢測
 - Immunocompromised
 - Requiring hospitalization
 - During period of influenza and COVID-19

- Molecular assay
- Antigen detection assay
- Others

- Low to moderate sensitivity
- High specificity

- Low to moderate sensitivity 50~70%
- High specificity

Antigen detection assay is alternative

Interpret with caution

- Interpret with caution
 - False negative result are common

- Interpret with caution
 - False negative result are common
- Not used for hospitalized

Antigen detection assay is alternative

Need molecular test recheck

- Need molecular test recheck
 - Negative Ag in high community influenza, and indication for confirmation

- Need molecular test recheck
 - Negative Ag in high community influenza, and indication for confirmation
 - Positive Ag in low community influenza

- Need molecular test recheck
 - Negative Ag in high community influenza, and indication for confirmation
 - Positive Ag in low community influenza
 - Recent exposure to pigs or poultry

- Molecular assay
- Antigen detection assay
- Others

Others

Viral culture

Others

- Viral culture
- Serological tests

Others

- Viral culture: public surveillance
- Serological tests

Others

- Viral culture: public surveillance
- Serological tests: not routine test

- 流感診斷工具

- Why to test
- Whom to test
- What to test
- How to test



-How to test

• 檢測時間點

How to test

- 檢測時間點
- 採哪裡

How to test

- 檢測時間點
- 採哪裡
- •用什麼採檢

- 檢測時間點

Within 4 days of symptom onset

• 首選為Nasopharynx

- 首選為Nasopharynx
- 其次是Nasal加上Throat swab

- 首選為Nasopharynx
- 其次是Nasal加上Throat swab
 - 若非要選一個,就選nasal swab

 Greater and prolonged respiration in lower respiratory tract

- Greater and prolonged respiration in lower respiratory tract
- Ventilated patients with negative upper respiratory tract
 - Endotracheal aspirate
 - Bronchoalveolar lavage fluid

-用什麼採

-用什麼採

Flocked swab

-用什麼採

Flocked swab



-Case



42 years old

法定傳染病通報送檢檢驗報告

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Influenza virus type B antigen Negative

-Case



42 years old

Rapid molecular test-Flu A Positive

Viral nucleic acid detection-influenza A virus(病毒核酸檢驗-A型流感病毒)
Positive (H1),請確認是否通報流感併發重症

-診斷工具-

- 當檢驗結果會影響後續決策或是要進行 流病調查時,才需要進行流感的檢測
- 流感診斷以核酸檢測為原則,若是用抗原檢測,則需要小心判讀
- 在病程早期進行檢測的陽性率較高,最好是取鼻咽的檢體

流感臨床診斷與檢驗

臨床表現

診斷工具

鑑別診斷

- Case



26 years old

- Rheumatoid arthritis

-Case



26 years old

Cough since last night

- Fever
- Headache
- Chest pain
- Nausea/vomiting, diarrhea

-Case

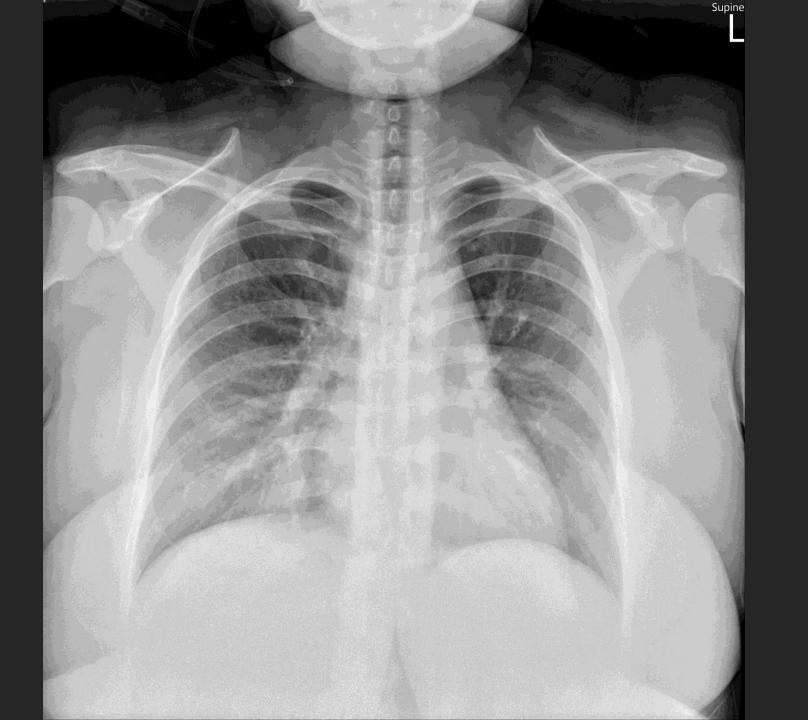


26 years old

Cough since last night

- Fever
- Headache
- Chest pain
- Nausea/vomiting, diarrhea

Dyspnea with desaturation



是什麼診斷?



Signs and symptom for pneumonia

- Signs and symptom for pneumonia
- Abnormal lung auscultation

- Signs and symptom for pneumonia
- Abnormal lung auscultation
- Persistent fever > 3~5 days

- Signs and symptom for pneumonia
- Abnormal lung auscultation
- Persistent fever > 3~5 days
- Fever after initial defervescence

- Signs and symptom for pneumonia
- Abnormal lung auscultation
- Persistent fever > 3~5 days
- Fever after initial defervescence
- Worsening symptom

Respiratory viral infection

- Respiratory viral infection
- Bacterial pneumonia

- Respiratory viral infection
- Bacterial pneumonia

- COVID-19
- Respiratory syncytial virus
- Common cold
- MERS-CoV
- Novel influenza A virus infection

COVID-19

Difficult to distinguish

COVID-19

- Difficult to distinguish
- More common in COVID-19
 - Fatigue
 - Diarrhea
 - Olfactory disorders
 - Taste disorders

COVID-19

Diagnosed by antigen / PCR

Respiratory syncytial virus

More common in children

Respiratory syncytial virus

- More common in children
- Also important in
 - Older adults
 - Immunocompromised

Common cold

- Several virus
 - Rhinovirus
 - Parainfluenza
 - Common cold coronavirus

Common cold

- Mild symptom than influenza
- Nasal congestion milder
- Low risk of complication

Common cold

Diagnosis by clinical manifestation

MERS-CoV

History of Arabian peninsula

MERS-CoV

- High risk of complication
 - Acute kidney injury
 - Disseminated intravascular coagulation

MERS-CoV

Diagnosis by PCR

Novel influenza A virus

 Symptom not distinguish from seasonal influenza A

Novel influenza A virus

- Exposure history
- Travel history

Novel influenza A virus

- Exposure to
 - Poultry
 - Pigs
 - Ill person with animal associated influenza

Novel influenza A virus

- Travel to region with local transmission
 - H7N9: China
 - H5N1: Asia, Middle east

Novel influenza A virus

Diagnosis by PCR

其它鑑別診斷

Bacterial pneumonia

- Complication of influenza
- Infection alone

其它鑑別診斷

Bacterial pneumonia

- Complication of influenza
 - Initial improvement
 - Relapse of fever and productive cough

其它鑑別診斷

Bacterial pneumonia

- Fever
- Dyspnea
- Cough
- Sputum production

-Case



26 years old

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Influenza virus type A antigen Positive
請確認是否通報流感併發重症
Influenza virus type B antigen Negative

SARS-CoV-2 Ag (公費快篩-有症狀/醫用) Positive

一鑑別診斷-

- 各種會造成呼吸道感染的病原體都要考慮
 - Common cold以臨床診斷就可以
 - MERS-CoV和新型A流則需要利用核酸作診斷
- •除了病毒,細菌性肺炎也是需要思考

-Take home message -

- 在只有流感流行時,臨床診斷的正確率可達80~90%
- 流感除了呼吸道症狀,全身性的症狀會特別明顯,像是發燒、 頭痛、痠痛等
- 在流感非流行期,有相關的曝露史,再進行流感相關檢測
- 流感的檢測以核酸為原則,使用抗原時要謹慎解讀
- 除了流感,呼吸道相關的感染症也要考慮。
- 適時地診斷出流感,才能及時給予適當的治療