

## Transplacental infections

classification	Timing of event	mechanism
<p><b><u>Congenital:</u></b> Mostly viral [ <b>TORCH</b> ]</p> <ul style="list-style-type: none"> <li>• <b>T</b>oxoplasmosis</li> <li>• <b>O</b>ther [ syphilis - parvo - VZV ]</li> <li>• <b>R</b>ubella</li> <li>• <b>CMV</b></li> <li>• <b>H</b>erpes [ Hepatitis - HIV ]</li> </ul> <p><b>Common findings:</b> majority <b>asymptomatic</b></p> <ul style="list-style-type: none"> <li>• Intrauterine growth retardation(IUGR)</li> <li>• Hepatosplenomegaly(HSM)</li> <li>• Thrombocytopenia</li> <li>• Microcephaly</li> </ul> <p><u>Preventative and therapeutic measures possible for some agents</u></p>	In utero	<p style="color: red;"><u>Trans placental</u></p> <p>Risk of IUI &amp; fetal damage:</p> <ul style="list-style-type: none"> <li>• Type of organism (teratogenic)</li> <li>• Type of maternal inf.(1<sup>o</sup>, Recurrent )</li> <li>• Time of inf .(1<sup>st</sup>, 2<sup>nd</sup> or 3<sup>rd</sup>)</li> </ul> <p>1<sup>o</sup> Maternal infection in the first half of pregnancy poses <u>the greatest risk</u> to the fetus</p>
<b><u>Perinatal</u></b>	During labor	Exposure to genital secretions and blood
<b><u>Neonatal</u></b>	After birth	Direct contact breast feeding or nosocomial exposure
<p><b>Serological DX:</b></p> <ul style="list-style-type: none"> <li>• <b>IgM</b> antibody [ absence of fetal IgM at birth does not exclude inf ]</li> <li>• Persistence of specific <b>IgG</b> antibody &gt;12 ms of age</li> </ul>		

### **Tocoplasma Gondii [ congenital obligate intracellular parasite ]**

<b>Forms</b>	<ul style="list-style-type: none"> <li>• <b>Oocyst:</b> shed in cat feces</li> <li>• <b>Tachyzoite:</b> rapidly dividing acute stage</li> <li>• <b>Bradyzoite:</b> slowly dividing chronic phase</li> </ul> <p style="color: red;"><b>If a person is Bradyzoite and there is a decrease in immunity they will go into the tachyzoite phase</b></p>
<b>Transmission</b>	<ul style="list-style-type: none"> <li>• Ingestion of Oocyst: from contaminated fingers / soil / water</li> <li>• ingestion of cyst : from undercooked meat</li> <li>• blood transfusion or organ transplant</li> </ul>
<b>Infection info</b>	<ul style="list-style-type: none"> <li>• most are due to 1ry maternal infection</li> <li>• Rarely from reactivation of latent inf</li> <li>• Most (70-90%) are <b>asymptomatic</b> at birth but are still at high risk of developing abnormalities: <ul style="list-style-type: none"> <li>○ <b>Classic triad :</b> Chorioretinitis / neurologic [ hydrocephalus - intracranial calcification ]</li> <li>○ <b>Other signs:</b> rash, HSM, jaundice, LAP, microcephaly, seizures, thrombocytopenia.</li> <li>○ <b>Abortion &amp; IVD</b></li> </ul> </li> </ul>
<b>Dx</b>	<ul style="list-style-type: none"> <li>• <b>Pregnant mother:</b> serology - IgM - IgG - IgG avidity - IgG seroconversion</li> <li>• <b>Infant:</b> <ul style="list-style-type: none"> <li>○ <b>Prenatal:</b> PCR - Culture - serial U.S</li> <li>○ <b>Postnatal:</b> <ul style="list-style-type: none"> <li>▪ serology [ IgG - IgM / ↑ IgG or persistently +ve &gt;12 ms ]</li> <li>▪ PCR</li> <li>▪ Culture</li> <li>▪ Evaluation of infant (ex, neuroimaging)</li> </ul> </li> </ul> </li> </ul>
<b>Tx &amp; prevention</b>	<ul style="list-style-type: none"> <li>• <b>Spiramycin. pyrimethamine &amp; sulfadiazine</b></li> <li>• avoid cat feces - wash hands / fruits / surfaces raw meat - cook all meats well</li> </ul>

## Parvovirus B19 [ congenital - Parvoviridae / non enveloped - icosahedral - ssDNA Virus ]

Epidemiology	<ul style="list-style-type: none"> <li>Worldwide distribution</li> <li>Humans are known hosts</li> <li>Transmission [ Respiratory route - Transplacental route - Blood transfusion ]</li> </ul>
Clinical	<ul style="list-style-type: none"> <li>Acquired: immunocompetent - immunocompromised --&gt; <b>Erythema infectiosum</b></li> <li>Congenital : Risk of congenital infection is greatest when inf occur in 1<sup>st</sup> 20 wks <ul style="list-style-type: none"> <li>Inf in the 1<sup>st</sup> trimester --&gt; IUD (Intrauterine death)</li> <li>Inf in the 2<sup>nd</sup> trimester --&gt; HF (Hydrops fetalis)</li> <li>Inf in the 3<sup>rd</sup> trimester --&gt; Lowest risk</li> </ul> </li> </ul> <p>Cause fetal loss through hydrops fetalis, severe anaemia, CHF, generalized oedema and fetal death</p>
Dx	<ul style="list-style-type: none"> <li><b>Pregnant mother:</b> Specific IgM - IgG seroconversion</li> <li><b>Prenatal :</b> Not grow in cell culture - PCR - U.S (hydrops)</li> </ul>
Tx	<ul style="list-style-type: none"> <li>Intrauterine transfusion</li> </ul>
prevention	<ul style="list-style-type: none"> <li>Hygiene practice - No vaccine (TRIAL)</li> </ul>


## Varicella zoster virus / VZV [congenital - Herpesviridae/ enveloped - icosahedral - dsDNA Virus ]

Transmission	<ul style="list-style-type: none"> <li>Respiratory &amp; Transplacental</li> </ul>
Clinical	<ul style="list-style-type: none"> <li><b>Acquired:</b> <ul style="list-style-type: none"> <li>Varicella / Chickenpox: 1<sup>o</sup> illness - <b>Generalized</b> vesicular rash</li> <li>Zoster / Shingles: Recurrent form - <b>Localized</b> Vesicular Rash</li> </ul> </li> <li><b>Congenital :</b> <ul style="list-style-type: none"> <li>VZV infection in Pregnancy: <ul style="list-style-type: none"> <li>Primary infection carries a greater risk of severe disease in particular pneumonia and its rare</li> </ul> </li> <li><b>Intrauterine infections</b> <ul style="list-style-type: none"> <li><b>congenital varicella syndrome:</b> 1st 20 ws of preg / CVS ~2% / skin scarring / limbs hypoplasia / CNS &amp; eye defects</li> <li><b>Neonatal varicella:</b> &lt;5 d of delivery --&gt; severe / &gt;5 d before delivery --&gt; mild</li> </ul> </li> </ul> </li> </ul>
Dx	<ul style="list-style-type: none"> <li><b>Pregnant mother:</b> <ul style="list-style-type: none"> <li><b>Direct sample:</b> <ul style="list-style-type: none"> <li>Vesicular fluid for virus isolation</li> <li>Cells scraping from the base of vesicles --&gt; IF test (Ag)</li> <li>DNA-VZV by PCR</li> </ul> </li> <li><b>Serology : IgM AB</b></li> </ul> </li> <li><b>Infant:</b> <ul style="list-style-type: none"> <li><b>Prenatal:</b> <ul style="list-style-type: none"> <li>VZV DNA in Fetal BI or AF or placenta villi</li> <li>VZV IgM in Fetal BI.</li> <li>U.S</li> </ul> </li> <li><b>Postnatal:</b> <ul style="list-style-type: none"> <li>VZV IgM</li> <li>virus isolation</li> <li>VZV DNA in VF or CSF ( CNS INF )</li> </ul> </li> </ul> </li> </ul>
Tx	<ul style="list-style-type: none"> <li><b>Acyclovir</b></li> </ul>
Prevention	<ul style="list-style-type: none"> <li><b>Post exposure: VZIG</b> <ul style="list-style-type: none"> <li>susceptible pregnant women have been exposed to VZV.</li> <li>infants whose mothers develop V &lt; 5 days of delivery or the first 2 days after delivery.</li> </ul> </li> <li><b>Pre exposure: live attenuated vaccine</b></li> </ul>

## Rubella virus [congenital - Togaviridae / enveloped - icosahedral - ssRNA Virus ]

Transmission	<ul style="list-style-type: none"> <li>• Respiratory &amp; Transplacental</li> </ul>
Pathogenesis	<ul style="list-style-type: none"> <li>• infection of nasopharynx and lungs of the mother --&gt; goes to Lymph nodes&amp; reticuloendothelial system --&gt; blood --&gt; if she is pregnant --&gt; penetration of placenta --&gt; infect fetus</li> <li>• Classic triad / CRS :             <ul style="list-style-type: none"> <li>○ <b>Cataracts - deafness - Cardiac problems (PDA)</b>- Neural defects - growth retardation - bone disease - HSM - <b>↓ PLT [ Blueberry muffin lesion ]</b></li> </ul> </li> </ul>
Clinical	<ul style="list-style-type: none"> <li>• <b>Acquired: Maculopapular rash (German measles)</b></li> <li>• <b>Congenital : [ Normal --&gt; Congenital Rubella Syndrom --&gt; IntraUterine Death ]</b></li> </ul> <p style="text-align: center; color: red;">Risk of acquiring congenital rubella infection varies and depends on gestational age of the fetus at the time of maternal infection. first 12 w --&gt; 70% risk</p>
Dx	<ul style="list-style-type: none"> <li>• <b>Pregnant mother:</b> serology [ Rubella specific IgM - IgG seroconversion ]</li> <li>• <b>Infant:</b> <ul style="list-style-type: none"> <li>○ Cell culture &amp; RT-PCR               <ul style="list-style-type: none"> <li>▪ (amniotic fluid &amp; chorionic villi) fetus</li> <li>▪ (nasal secretion , throat, urine &amp; blood) newborn</li> </ul> </li> <li>○ Serological diagnosis               <ul style="list-style-type: none"> <li>▪ Rubella specific IgM</li> <li>▪ Persistence &amp; rising titres of anti-rubella</li> <li>▪ IgG in the infants serum beyond 9-12 ms of age</li> </ul> </li> </ul> </li> </ul>
Prevention	<ul style="list-style-type: none"> <li>• <b>Routine antenatal screening:</b> <ul style="list-style-type: none"> <li>○ Rubella specific IgG</li> <li>○ Non-immune women --&gt; vaccination ( avoid pregnancy for 3 months).</li> </ul> </li> <li>• <b>vaccination :</b> <ul style="list-style-type: none"> <li>○ before or after pregnancy but not during pregnancy.</li> </ul> </li> </ul>

## CMV [congenital - Herpesviridae / enveloped - icosahedral - dsDNA Virus ]

Transmission	<ul style="list-style-type: none"> <li>• Horizontal :             <ul style="list-style-type: none"> <li>○ Children -&gt; saliva</li> <li>○ later in life --&gt; sexual - Blood - organ transplant</li> </ul> </li> <li>• Vertical :             <ul style="list-style-type: none"> <li>○ 1ry --&gt; 40%</li> <li>○ recurrent --&gt; 1%</li> </ul> </li> </ul> <p>* latent form --&gt; reactivation --&gt; recurrent inf          * Transplacental - during birth [ infected canal ] - breast milk          * mostly asymptomatic</p>	 <p style="border: 1px solid black; padding: 2px; display: inline-block;">Blueberry muffin</p>
Cytomegalic inclusion disease	<ul style="list-style-type: none"> <li>• CNS abnormalities [ microcephaly, periventricular calcification ]</li> <li>• Eye - chorioretinitis <span style="float: right;">* Ear - sensorineural deafness</span></li> <li>• Liver – HSM and jaundice. <span style="float: right;">*Lung - pneumonitis</span></li> <li>• Heart - myocarditis <span style="float: right;">*Thrombocytopenic purpura</span></li> </ul>	
Dx	<ul style="list-style-type: none"> <li>• <b>Pregnant mother:</b> serology [CMV IgM - CMV IgG - IgG Avidity]</li> <li>• <b>Infant:</b> <ul style="list-style-type: none"> <li>○ Prenatal : PCR - Culture - CMV IgM - <b>U.S</b></li> <li>○ Post natal: by <i>isolating CMV or detection of its genome</i> in <b>first 3 wks</b> of life               <ul style="list-style-type: none"> <li>▪ <b>Body fluid :</b> urine, saliva, blood By [ Standard tube culture method - Shell vial assay-PCR ]</li> <li>▪ <b>Histology:</b> inclusion bodies [ owl's eyes ]</li> <li>▪ <b>serology:</b> CMV IgM</li> </ul> </li> </ul> </li> </ul>	
Tx & prevention	<ul style="list-style-type: none"> <li>• Symptomatic infant --&gt; Ganciclovir <span style="float: right;">* asymptomatic --&gt; not recommended</span></li> <li>• Education [ Hygiene - hand washing] - Vaccine [ not available / trial ]</li> </ul>	