Vaccination In Renal Transplantation



General Consideration

<u>Introduction</u>

Vaccines are an important preventative tool and offer protection against infection

Introduction

At increased risk of infectious complications of vaccine-preventable diseases

Diminished response to many vaccines in organ failure

Transplant candidates and recipients

Full complement of recommended vaccinations prior to transplantation

Immunization early in the course of kidney failure

Pre-Transplant Vaccination

- ➤ Ideally, KT recipients should be vaccinated as early as possible as the response to vaccines is diminished in end-organ failure and in states of immunosuppression
- ➤ It is recommended to vaccinate patients with CKD, not requiring dialysis, so that they can mount an optimal immunological response

World J Transplantation 2019 January 16; 9(1): 1-13 Clinical Transplantation. 2019;33:e13563

Pre-Transplant Vaccination

Vaccination status should be reviewed at the first transplant clinic visit

 For patients who are incompletely or unvaccinated prior to transplant, consultation with an infectious disease specialist is recommended

Clinical Transplantation. 2019;33:e13563

If feasible



Vaccines should be administered prior to planned immunosuppression

- Live vaccines:
 - —≥4 w prior to transplantation
 - Avoided within 2 w of transplantation



After administration of live viral vaccines: □Viral replication and immunologic response <3 w ■Vaccination ≥4 w prior to transplant is safe

Live Vaccine

- Live vaccine may interfere with the reading of tuberculin skin test and IGRA
- Therefore, the TST should be performed simultaneously with live vaccine or delayed by at least 28 d

 Inactivated vaccines should be administered

≥2 w

prior to transplantation



In solid-organ recipients receiving immunosuppression, the immune system will not be able to mount a response as effective as in normal subjects

Most immunosuppressive regimens after solid-organ transplant

Steroids

Calcineurin inhibitors

Cyclosporin and Tacrolimus

Both T and B cell responses are impaired through blockage of cellular proliferation after antigen stimulation as well as inhibition of cytokine production necessary for such stimulation

Potent cytokine inhibitors: Interleukin-1, 2, 6, TNF and gamma interferon

Block antigen-induced T-cell proliferation

Corticosteroids

Immunosuppression with steroids alone does not seem to completely impair the immune response to vaccine administration

Calcineurin inhibitors

Directly inhibit interleukin-2 dependent T-cell proliferation

Blocking interleukin-4 and 5 production by T cells with inhibitory effect on B-cell function and antibody production

- ➤ Interfere with purine synthesis
- ➤ Blocking both T- and B-cell proliferation

Azathioprine and Mycophenolate mofetil

Post-transplantation hypogammaglobulinemia

diminished protective response to pneumococcus and Td vaccine in heart, **kidney** and lung transplant



The combination of these mechanisms leads to significant impairment of the entire immunologic cascade following antigen presentation to immune cells

- The production of new memory cells and the survival of memory cells acquired prior to transplantation is critical to an effective response to vaccines
- The effect of immunosuppression on immune memory cells is not completely understood and the specific life span of memory T cells have not been determined in these patients

Timing of Vaccination Post Transplant

 If a vaccination series is initiated pre-transplant but not completed prior to transplantation, continuation can occur in the post-transplant period



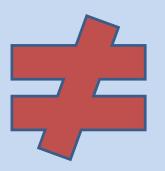
Timing of Vaccination Post Transplant

- Standard age-appropriate inactivated vaccine series should be administered 2 - 6 months after SOT(3-6 m)
- IIV can be administered ≥1 month after transplant during a community influenza outbreak

Post Transplant Vaccination

AST 2019

Vaccination during active treatment for rejection should be avoided



IDSA 2013

Vaccination should not be withheld because of concern about transplant organ rejection

Post Transplant Vaccination

 MMR vaccine and VAR should generally not be administered to SOT recipients

Except for varicella:

- ✓ In children without evidence of immunity who are renal or liver transplant recipients
- ✓ Are receiving minimal or no immunosuppression
- ✓ Have no recent graft rejection

Serologic Response in KT

- KTs are on life-long immunosuppression
- After vaccination
 - Lower rates of seroconversion
 - Lower mean antibody titers
 - Waning of protective immunity over shorter period

as compared to general population

Serologic Response in KT

Seroconversion should be documented by serologic assays where available

A minimum of 4 weeks should elapse between vaccine administration and evaluation for seroconversion

Vaccination before & after KT

Vaccine	Recommended before KT	Recommended after KT
HBV	Yes	Yes
Influenza	Yes	Yes
Tdap/ Td	Yes	Yes
Pneumococcal vaccine	Yes	Yes
Haemophilus influenza type B	Yes	Yes

Vaccination before & after KT

Vaccine	Recommended before KT	Recommended after KT
HPV	Yes	Yes
Varicella Zoster	Yes	No
Rotavirus	Yes	No
MMR	Yes	No
IPV/OPV	Yes	Yes for IPV No for OPV

Evaluation for Serologic Response

- HBV> 10 mIU/ml
- Tetanus
- H.influenzae vaccine-induced anticapsular
 Ab(polyribosylribitolphosphate) >0.15 mg/L
- MMR



<u>Introduction</u>

 Every effort should be made to ensure that transplant candidates and their household members have completed the full complement of recommended vaccinations prior to transplantation

Individuals who live in a household with immunocompromised patients can receive inactivated vaccines

- Possible transmission through close contact
 - Live intranasal influenza
 - Oral polio
 - Smallpox vaccines

should be deferred in HCWs and household members of transplant recipients when possible

Age ≥6 months should receive influenza vaccine annually

They should receive either:

(a) IIV

OR

(b) LAIV provided they are healthy, not pregnant, and aged 2–49 y

If LAIV administered in this situation

Contact with the immunocompromised patient should be avoided for 7 d?

 OPV should not be administered to individuals who live in a household with immunocompromised patients



Other live attenuated vaccines

- Varicella/Zoster
- MMR
- Rotavirus in infants aged 2–7 m
 are unlikely to result in transmission from normal host to immunocompromised host

Immunocompromised patients should avoid contact with persons who develop skin lesions after receipt VAR or ZOS until the lesions clear

 Transplant recipients caring for infants who have been given rotavirus vaccine should defer diaper changing for 2-4 weeks and otherwise enact scrupulous hand hygiene

Indicated Vaccines

Inactivated vaccines	Influenza	Live Vaccine
Td/Tdap	IIV	MMR
Pneumococcal HPV	LAIV	Zoster
		Varicella
пру		Rotavirus

Pet Immunization

- Pets should also be fully immunized
- There is little or no risk of transmission following immunization of pets with live vaccines





References

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